

10/577531

Connecting via Winsock to STN

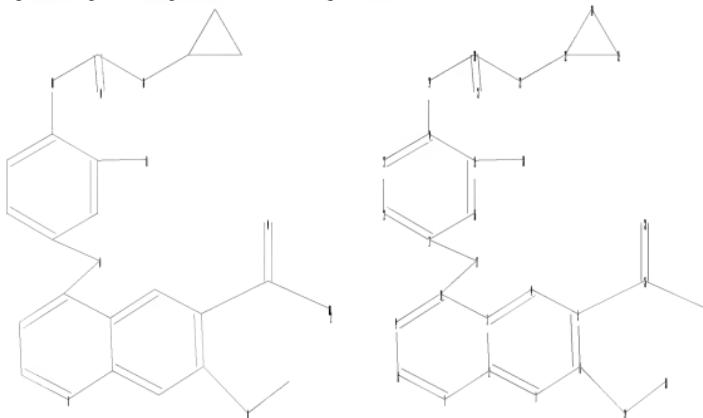
Welcome to STN International! Enter x:x

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FILE 'HOME' ENTERED AT 14:11:10 ON 10 SEP 2008

=> file reg

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Uploading C:\Program Files\Stnexp\Queries\10577531.str



chain nodes :
17 18 19 20 21 23 26 27 28 29 30
ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 22 24 25
chain bonds :
5-26 6-29 10-17 11-17 14-19 15-18 19-20 20-21 20-23 21-22 26-27 26-28
29-30
ring bonds :
1-2 1-6 2-3 2-7 3-4 3-10 4-5 5-6 7-8 8-9 9-10 11-12 11-16 12-13 13-14
14-15 15-16 22-24 22-25 24-25
exact/norm bonds :
6-29 10-17 11-17 14-19 19-20 20-21 20-23 21-22 22-24 22-25 24-25 26-27
26-28 29-30
exact bonds :
5-26 15-18
normalized bonds :

10/577531

1-2 1-6 2-3 2-7 3-4 3-10 4-5 5-6 7-8 8-9 9-10 11-12 11-16 12-13 13-14
14-15 15-16

Match level :

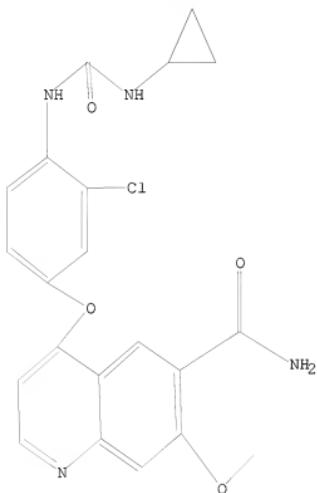
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:CLASS 18:CLASS 19:CLASS
20:CLASS 21:CLASS 22:Atom 23:CLASS 24:Atom 25:Atom 26:CLASS 27:CLASS
28:CLASS 29:CLASS 30:CLASS

L1 STRUCTURE UPLOADED

=> d 11

L1 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 11 full

L3 35 SEA SSS FUL L1

=> file ca

=> s 13
L4 22 L3

=> d ibib abs hitstr 1-22

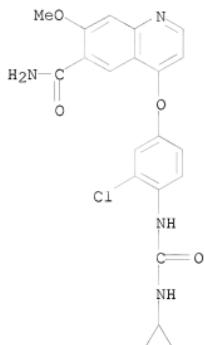
L4 ANSWER 1 OF 22 CA COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 149:239320 CA
TITLE: Composition for treatment of undifferentiated-type of
gastric cancer
INVENTOR(S): Yamamoto, Yuji
PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
SOURCE: PCT Int. Appl., 221pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2008093855 | A1 | 20080807 | WO 2008-JP51697 | 20080128 |
| W: AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BS, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, NO, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |

PRIORITY APPLN. INFO.: US 2007-887006P P 20070129
AB Disclosed are: a therapeutic agent, a kit and a treatment method for
undifferentiated-type of gastric cancer; and a pharmaceutical composition, a
kit and a treatment method which are more effective on a living body
having at least one cell selected from the group consisting of a cell
over-expressing FGFR2 and a cell expressing a FGFR2 mutant. A combination
of a FGFR2 inhibitor and a therapeutic substance for gastric cancer is
more effective on undifferentiated-type of gastric cancer. The
combination of a FGFR2 inhibitor and a therapeutic substance for gastric
cancer is more effective on a living body having at least one cell
selected from the group consisting of a cell over-expressing FGFR2 and a
cell expressing a FGFR2 mutant. For example, the synergistic effect of
combination of 4-(3-chloro-4-[cyclopropylaminocarbonyl]aminophenoxy)-7-
methoxy-6-quinolinecarboxamide and irinotecan hydrochloride in HSC-30
human gastric carcinoma cell-bearing mice was examined
IT 417716-92-8, 4-(3-Chloro-4-[cyclopropylaminocarbonyl]aminophenoxy)-
7-methoxy-6-quinolinecarboxamide 857890-39-2
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
composition for treatment of undifferentiated-type of gastric cancer
containing
quinoline derivs. in combination with antitumor agent or FGFR2
inhibitor)

10/577531

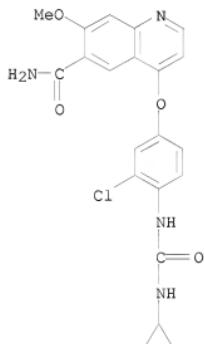
RN 417716-92-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



RN 857890-39-2 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2
 CMF C H4 O3 S



REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 149:167954 CA
 TITLE: Composition for treatment of pancreatic cancer
 INVENTOR(S): Yamamoto, Yuji
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 126pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------------------------|--------------------------|
| WO 2008088008 | A1 | 20080724 | WO 2008-JP51024 | 20080118 |
| W: AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, NO, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| PRIORITY APPLN. INFO.: | | | US 2007-885733P US 2007-887010P | P 20070119 P 20070129 |

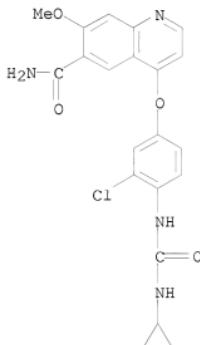
OTHER SOURCE(S): MARPAT 149:167954
 AB Disclosed are a pharmaceutical composition having excellent antitumor activity, and a method for treating a cancer. Specifically, excellent antitumor activity is achieved when 4-(3-chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide (A) or an analogous compound thereof, a pharmacol. acceptable salt thereof or a solvate of any of them is used in combination with gemcitabine or erlotinib, a pharmacol. acceptable salt thereof or a solvate of any of them. For example, the effect of combination of a compound A 3 mg/kg and gemcitabine hydrochloride 200 mg/kg on AsPC-1 human pancreatic cancer cell-bearing mice was examined 417716-92-8, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide 417717-05-6,
 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-methoxyethoxy)-6-quinolinecarboxamide 417717-07-8, 4-(3-Chloro-4-

(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-(4-morpholino)ethoxy)-6-quinolinecarboxamide 417717-10-3, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-hydroxyethoxy)-6-quinolinecarboxamide 417717-15-8, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-((2S)-2,3-dihydroxypropyl)oxy-6-quinolinecarboxamide 417719-50-7, 4-(3-Chloro-4-(*cis*-2-fluoro-cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide 417719-56-3, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-ethoxyethoxy)-6-quinolinecarboxamide 417719-77-8, 4-(3-Chloro-4-(((cyclopropylamino)carbonyl)amino)phenoxy)-7-((2R)-2-hydroxy-3-(1-pyrrolidino)propoxy)-6-quinolinecarboxamide 857890-39-2

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(pharmaceutical compns. containing urea derivs. in combination with gemcitabine or erlotinib)

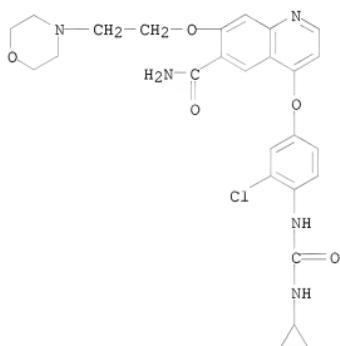
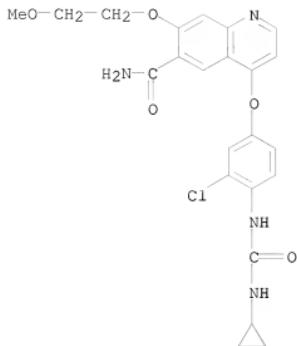
RN 417716-92-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)

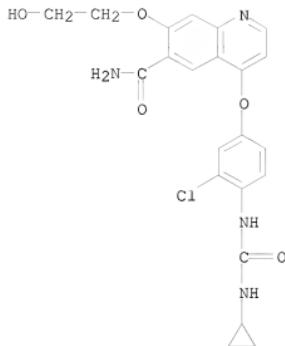


RN 417717-05-6 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)

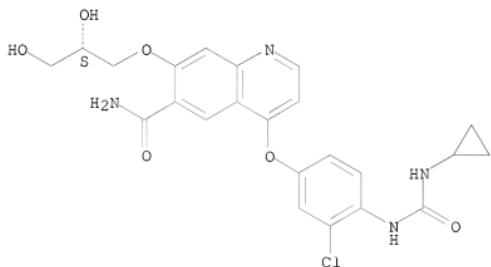


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



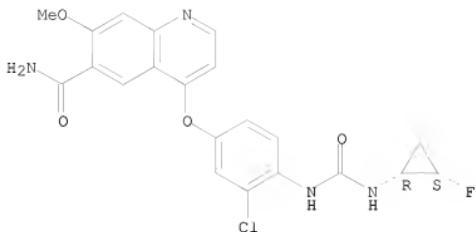
RN 417717-15-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



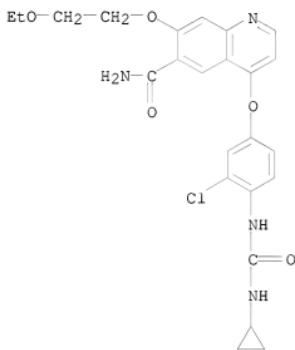
RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.



RN 417719-56-3 CA

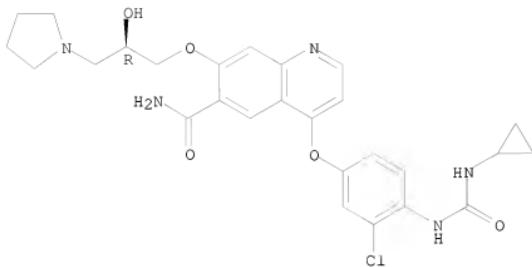
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA

CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[{(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy}-] (CA INDEX NAME)

Absolute stereochemistry.



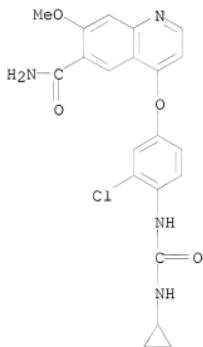
RN 857890-39-2 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

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CRN 417716-92-8

CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2

CMF C H4 O3 S

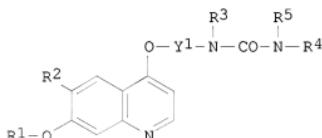


REFERENCE COUNT: 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 3 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESION NUMBER: 148:323091 CA
 TITLE: Antitumor agent for undifferentiated gastric cancer
 INVENTOR(S): Yamamoto, Yuji; Matsushima, Tomohiro; Tsuruoka,
 Akihiko; Obaishi, Hiroshi; Nakagawa, Takayuki
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 138pp.
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2008026748 | A1 | 20080306 | WO 2007-JP67088 | 20070827 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |

PRIORITY APPLN. INFO.: JP 2006-230816 A 20060828
 OTHER SOURCE(S): MARPAT 148:323091
 GI



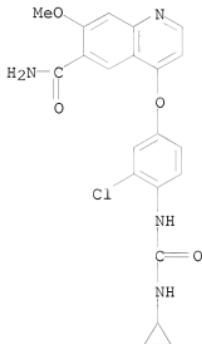
AB A compound represented by the general formula (I), a pharmacol. acceptable salt thereof, or a solvate of the compound or the salt can exert its effect

more effectively on undifferentiated gastric cancer, and can also exerts its effect more effectively on a living body having at least one member selected from the group consisting of a cell over-expressing FGFR2 and a cell expressing mutant FGFR2.

IT 417716-92-8P, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (quinolinylurea analogs as antitumor agents for undifferentiated gastric cancer)

RN 417716-92-8 CA

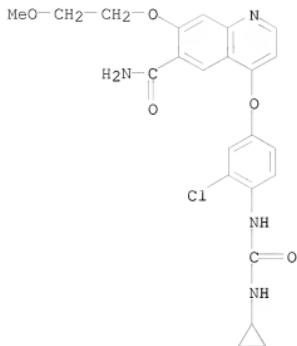
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



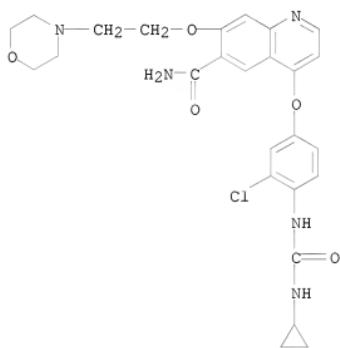
IT 417717-05-6, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-methoxyethoxy)-6-quinolinecarboxamide 417717-07-8
 417717-10-3 417717-15-8, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-((2S)-2,3-dihydroxypropyl)oxy-6-quinolinecarboxamide 417719-50-7 417719-56-3
 417719-77-8
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (quinolinylurea analogs as antitumor agents for undifferentiated gastric cancer)

RN 417717-05-6 CA

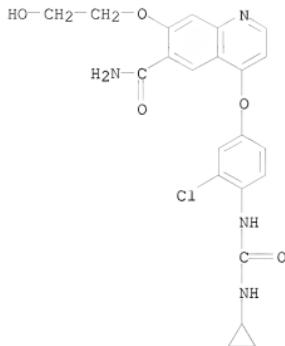
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-07-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

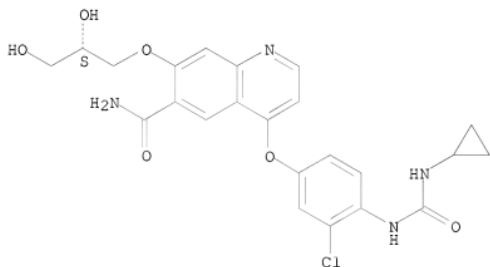


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



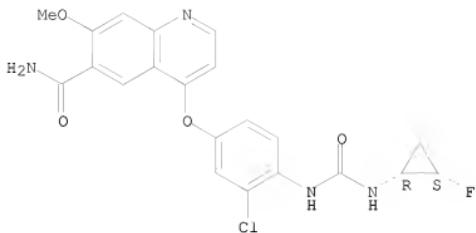
RN 417717-15-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



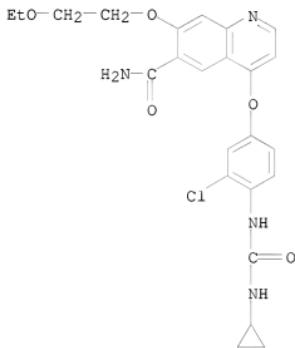
RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.



RN 417719-56-3 CA

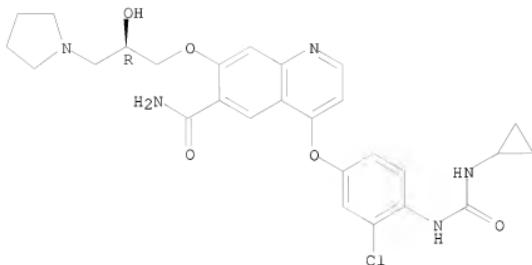
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA

CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[{(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy}-] (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 22 CA COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 148:253561 CA

TITLE: E7080, a novel inhibitor that targets multiple kinases, has potent antitumor activities against stem cell factor producing human small cell lung cancer H146, based on angiogenesis inhibition

AUTHOR(S): Matsui, Junji; Yamamoto, Yuji; Funahashi, Yasuhiro; Tsuruoka, Akihiko; Watanabe, Tatsuo; Wakabayashi, Toshiaki; Uenaka, Toshimitsu; Asada, Makoto

CORPORATE SOURCE: Tsukuba Research Laboratories, Tsukuba, Ibaraki, 300-2635, Japan

SOURCE: International Journal of Cancer (2007), Volume Date 2008, 122(3), 664-671
CODEN: IJCNAW; ISSN: 0020-7136

PUBLISHER: Wiley-Liss, Inc.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB E7080 is an orally active inhibitor of multiple receptor tyrosine kinases including VEGF, FGF and SCF receptors. In this study, we show the inhibitory activity of E7080 against SCF-induced angiogenesis in vitro and tumor growth of SCF-producing human small cell lung carcinoma H146 cells in vivo. E7080 inhibits SCF-driven tube formation of HUVEC, which express SCF receptor, KIT at the IC₅₀ value of 5.2 nM and it was almost identical for VEGF-driven one (IC₅₀ = 5.1 nM). To assess the role of SCF/KIT signaling in tumor angiogenesis, we evaluated the effect of imatinib, a selective KIT kinase inhibitor, on tumor growth of H146 cells in nude mice. Imatinib did not show the potent antitumor activity in vitro (IC₅₀ = 2,200 nM), because H146 cells did not express KIT. However, oral administration of imatinib at 160 mg/kg clearly slowed tumor growth of H146 cells in nude mice, accompanied by decreased microvessel density. Oral administration of E7080 inhibited tumor growth of H146 cells at doses of 30 and 100 mg/kg in a dose-dependent manner and caused tumor regression at 100 mg/kg. While anti-VEGF antibody also slowed tumor growth, it did not cause tumor regression. These results indicate that KIT signaling has a role in tumor angiogenesis of SCF-producing H146 cells, and E7080 causes regression of H146 tumors as a result of antiangiogenic activity mediated

by inhibition of both KIT and VEGF receptor signaling. E7080 may provide therapeutic benefits in the treatment of SCF-producing tumors.

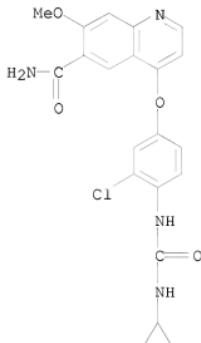
IT 417716-92-8

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(E 7080; E7080, a novel inhibitor that targets multiple kinases, has potent antitumor activities against stem cell factor producing human small cell lung cancer H146, based on angiogenesis inhibition)

RN 417716-92-8 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



REFERENCE COUNT: 38 THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 5 OF 22 CA COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 148:113266 CA

TITLE: Therapeutic agent for liver fibrosis

INVENTOR(S): Yokohama, Hiromitsu; Matsuoka, Toshiyuki

PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan

SOURCE: PCT Int. Appl., 82pp.

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|---|----------|-----------------|----------|
| WO 2008001956 | A1 | 20080103 | WO 2007-JP63525 | 20070629 |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, | | | |

TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW
 RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
 IS, IT, LT, LU, LV, MC, MT, NL, PL, PT, RO, SE, SI, SK, TR, BF,
 BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW,
 GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
 BY, KG, KZ, MD, RU, TJ, TM

PRIORITY APPLN. INFO.: US 2006-817872P P 20060629

OTHER SOURCE(S): MARPAT 148:113266

AB The object is to provide a therapeutic agent for liver fibrosis and a method for treatment of liver fibrosis. 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide or an analog thereof can prevent the fibrillation in the liver, and therefore can be used as a therapeutic agent for liver fibrosis or in the method for treatment of liver fibrosis.

IT 417716-92-8, 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-methoxy-6-quinolinecarboxamide 417717-05-6, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-methoxyethoxy)-6-quinolinecarboxamide 417717-07-8 417717-10-3 417717-15-8 417719-50-7 417719-56-3 417719-77-8

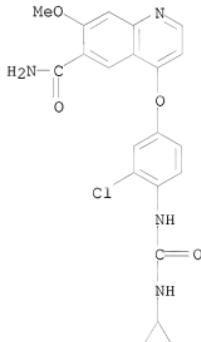
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide analogs as therapeutic agents for liver fibrosis)

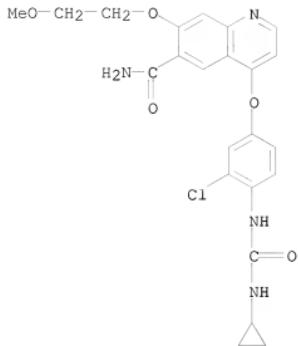
RN 417716-92-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)

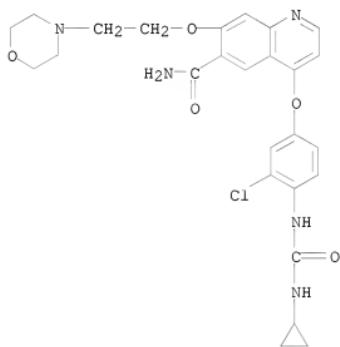


RN 417717-05-6 CA

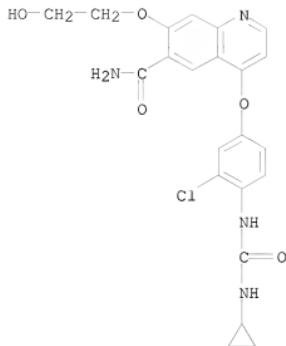
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-07-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

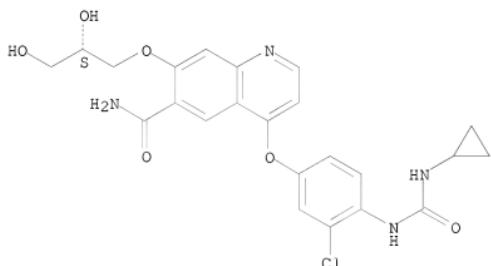


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



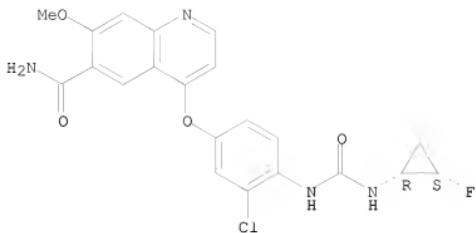
RN 417717-15-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



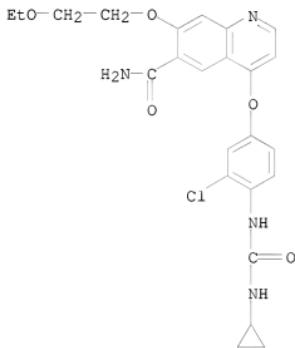
RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.



RN 417719-56-3 CA

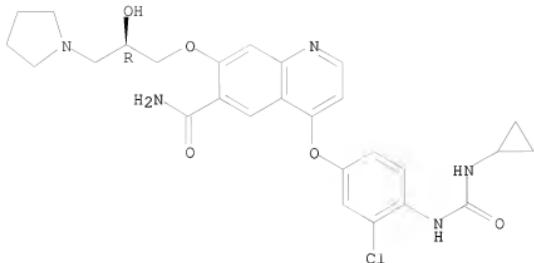
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 6 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 148:24395 CA
 TITLE: Antitumor agent for thyroid cancer containing RET kinase inhibitors
 INVENTOR(S): Matsui, Junji
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 140pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2007136103 | A1 | 20071129 | WO 2007-JP60560 | 20070517 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |

PRIORITY APPLN. INFO.: US 2006-747570P P 20060518

OTHER SOURCE(S): MARPAT 148:24395
 AB It is intended to provide a pharmaceutical composition exhibiting an effect more effectively on at least one disease selected from the group consisting of multiple endocrine neoplasia type IIA, multiple endocrine neoplasia type IIB, familial medullary thyroid carcinoma, thyroid cancer, papillary thyroid carcinoma, sporadic medullary thyroid carcinoma, Hirschsprung's disease, pheochromocytoma, parathyroid hyperplasia and

gastrointestinal mucosal neuroma; and a therapeutic method for the same. A compound 4-(3-chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide (I) and an analog thereof can exhibit an effect more effectively on at least one disease selected from the group consisting of multiple endocrine neoplasia type IIA, multiple endocrine neoplasia type IIB, familial medullary thyroid carcinoma, thyroid cancer, papillary thyroid carcinoma, sporadic medullary thyroid carcinoma, Hirschsprung's disease, pheochromocytoma, parathyroid hyperplasia and gastrointestinal mucosal neuroma. Usage of the RET kinase inhibitor for production of remedy for the diseases listed above, and a pharmaceutical composition containing the

RET

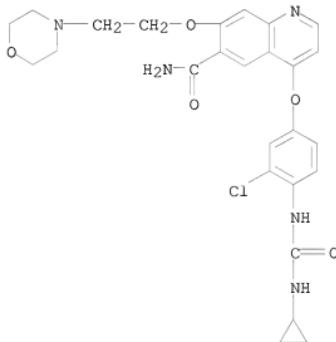
kinase inhibitor for treatment of biol. body including mutant RET protein, and method for prediction of sensitivity to RET kinase inhibitors through intracellular mutant RET protein as an indicator are also disclosed. For example, the inhibitory effect of I on RET kinase in human thyroid carcinoma cells (TT cells) was examined. Also, a coated tablet containing I methanesulfonate was formulated.

IT 417717-07-8

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-[2-(4-morpholino)ethoxy-6-quinolinecarboxamide; antitumor agent for thyroid cancer containing RET kinase inhibitors)

RN 417717-07-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)



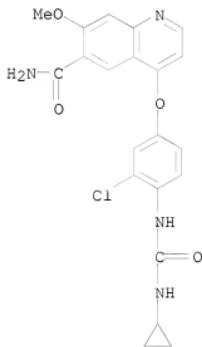
IT 417716-92-8 417717-05-6 417717-10-3,
 4-[3-Chloro-4-[(cyclopropylaminocarbonyl)amino]phenoxy-7-(2-hydroxyethoxy)-6-quinolinecarboxamide 417717-15-8, 4-[3-Chloro-4-[(cyclopropylaminocarbonyl)amino]phenoxy-7-[(2S)-2,3-dihydroxypropyl]oxy-6-quinolinecarboxamide 417719-50-7, 4-[3-Chloro-4-(cis-2-fluoro-cyclopropylaminocarbonyl)aminophenoxy]-7-methoxy-6-quinolinecarboxamide 417719-56-3, 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-(2-ethoxyethoxy)-6-quinolinecarboxamide 417719-77-8, 4-[3-Chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-

hydroxy-3-(1-pyrrolidino)propoxy]-6-quinolinecarboxamide
857890-39-2

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
(antitumor agent for thyroid cancer containing RET kinase inhibitors)

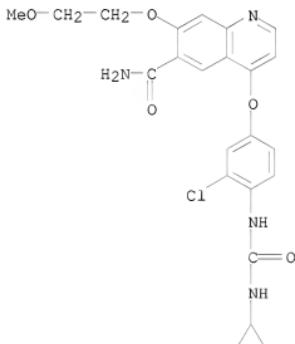
RN 417716-92-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
heoxy-7-methoxy- (CA INDEX NAME)



RN 417717-05-6 CA

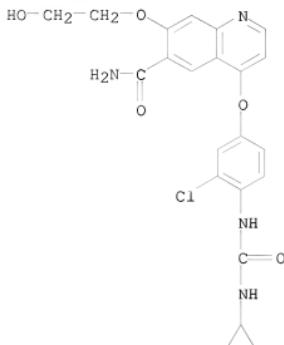
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
heoxy-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-10-3 CA

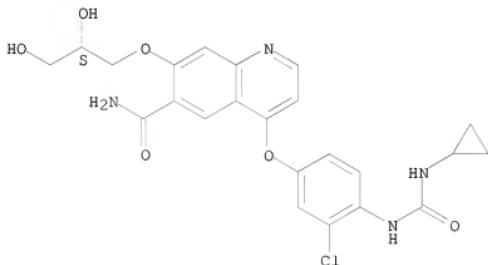
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p

heenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



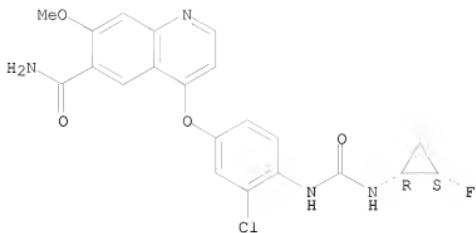
RN 417717-15-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



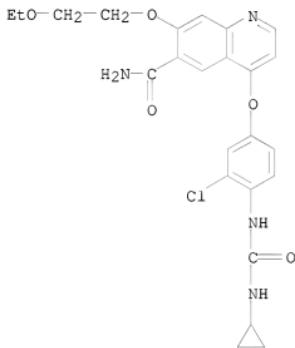
RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.



RN 417719-56-3 CA

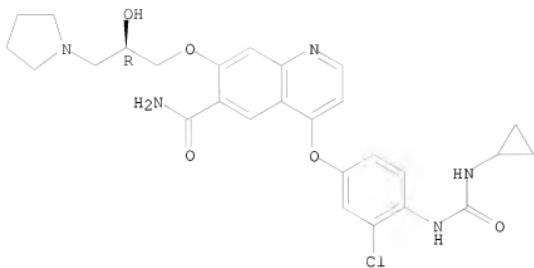
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA

CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

Absolute stereochemistry.



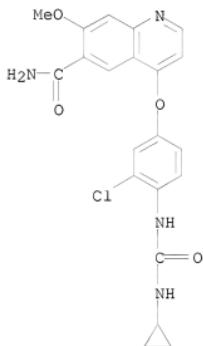
RN 857890-39-2 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8

CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2

CMF C H4 O3 S



L4 ANSWER 7 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 147:235192 CA
 TITLE: Preparation of urea derivatives containing nitrogenous aromatic ring compounds as inhibitors of angiogenesis
 INVENTOR(S): Funahashi, Yasuhiro; Tsuruoka, Akihiko; Matsukura, Masayuki; Haneda, Toru; Fukuda, Yoshio; Kamata, Junichi; Takahashi, Keiko; Matsushima, Tomohiro; Miyazaki, Kazuki; Nomoto, Ken-Ichi; Watanabe, Tatsuo; Obaishi, Hiroshi; Yamaguchi, Atsumi; Suzuki, Sachie; Nakamura, Katsuji; Mimura, Fusayo; Yamamoto, Yuji; Matsui, Junji; Matsui, Kenji; Yoshioka, Takako; Suzuki, Yasuyuki; Arimoto, Itaru
 PATENT ASSIGNEE(S): Eisai Co., Ltd, Japan
 SOURCE: U.S., 458pp., Cont.-in-part of Appl. No. PCT/JP01/09221.
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|----------|
| US 7253286 | B2 | 20070807 | US 2003-420466 | 20030418 |
| US 20040053908 | A1 | 20040318 | | |
| WO 2002032872 | A1 | 20020425 | WO 2001-JP9221 | 20011019 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1506962 | A2 | 20050216 | EP 2004-25700 | 20011019 |
| EP 1506962 | A3 | 20050302 | | |
| EP 1506962 | B1 | 20080702 | | |
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| EP 1777218 | A1 | 20070425 | EP 2006-23078 | 20011019 |
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| CN 101024627 | A | 20070829 | CN 2007-10007096 | 20011019 |
| CN 101029022 | A | 20070905 | CN 2007-10007097 | 20011019 |
| ES 2282299 | T3 | 20071016 | ES 2001-976786 | 20011019 |
| ZA 2003003567 | A | 20040810 | ZA 2003-3567 | 20030508 |
| JP 2005272474 | A | 20051006 | JP 2005-124034 | 20050421 |

| | | | | |
|----------------|----|----------|----------------|----------|
| US 20060247259 | A1 | 20061102 | US 2005-293785 | 20051202 |
| US 20060160832 | A1 | 20060720 | US 2006-347749 | 20060203 |
| AU 2006203099 | A1 | 20060810 | AU 2006-203099 | 20060719 |
| AU 2006236039 | A1 | 20061207 | AU 2006-236039 | 20061116 |
| AU 2006236039 | B2 | 20080522 | | |

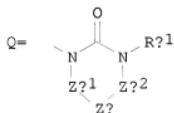
PRIORITY APPLN. INFO.:

| | |
|----------------|-------------|
| JP 2000-320420 | A 20001020 |
| JP 2000-386195 | A 20001220 |
| JP 2001-46685 | A 20010222 |
| WO 2001-JP9221 | A2 20011019 |
| AU 2001-295986 | A3 20011019 |
| AU 2001-95986 | TO 20011019 |
| CN 2001-819710 | A3 20011019 |
| EP 2001-976786 | A3 20011019 |
| JP 2002-536056 | A3 20011019 |
| US 2003-420466 | A3 20030418 |
| US 2005-293785 | A1 20051202 |

OTHER SOURCE(S):

MARPAT 147:235192

GI



AB N-aryl or N-heteroarylurea derivs. represented by the general formula Ag-Xg-Yg-Tg1 or salts thereof, or hydrates of both (wherein Ag = (un)substituted C6-14 aryl or 5- to 14-membered heterocyclic group; Xg = single bond, O, S, Cl-6 alkylene, SO, SO₂, (un)substituted NH; Yg = (un)substituted C6-14 aryl, 5- to 14-membered heterocyclic group, Cl-8 alkyl, C3-8 alicyclic hydrocarbyl, C6-14 aryl-Cl-6 alkyl, 5- to 14-membered heteroaryl-Cl-6 alkyl, (CH₂)_nSO₂ (g = 1-8), (CH₂)_{fa}CH:CH(CH₂)_{fb} (fa, fb = 0, 1, 2, 3), etc.; and Tg1 = a group of the general formula -Eg-CO-NRg1(Zg) or Q; wherein Eg = a single bond, (un)substituted NH; Rg1 = H, (un)substituted Cl-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, C3-8 aliphatic hydrocarbyl, etc.; Zg = Cl-8 alkyl, C3-8 alicyclic hydrocarbyl, C6-14 aryl, etc.; Zg1, Zg2 = (a) a single bond, (b) Cl-6 alkylene optionally having ≥1 atoms selected from O, S, and N in the middle or the terminus of the chain and optionally substituted with oxo, (c) (un)substituted C2-6 alkenyl] are prepared. These compds. are also inhibitors of vascular endothelial growth factor receptor kinase (VEGFR2 kinase) and are useful as antitumor agents against hemangioma, pancreatic cancer, stomach cancer, colon cancer, breast cancer, prostate cancer, lung cancer, brain tumor, leukemia, or ovarian cancer, as cancer metastasis inhibitors, and for the treatment of retina neovascularization, diabetic retinopathy, atherosclerosis, or inflammatory diseases such as osteoarthritis, rheumatoid arthritis, psoriasis, or delayed hypersensitivity. Thus, to solution of 334 mg 4-[6-(4-benzyloxyphenyl)-7-(2-trimethylsilylthoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy]-2-chlorophenylamine in 4 mL DMF were added 0.066 mL pyridine and 0.102 mL Ph chlorocarbonate and stirred at room temperature for 2.5 h to give 330 mg N-[4-[6-(4-benzyloxyphenyl)-7-(2-trimethylsilylthoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy]-2-chlorophenyl]-N'-cyclopropylurea which

(260 mg) was hydrogenolyzed over platinum oxide in ethanol overnight to give 160 mg N-[4-(6-(4-hydroxyphenyl)-7-(2-trimethylsilylethoxymethyl)-7H-pyrido[2,3-d]pyrimidin-4-yloxy)-2-chlorophenyl]-N'-cyclopropylurea (I). I showed IC₅₀ of 0.02 nM for inhibiting the vascular endothelial growth factor (VEGF)-stimulated sandwich tube formation in vascular endothelial cell.

IT 417717-12-5P 417717-13-6P

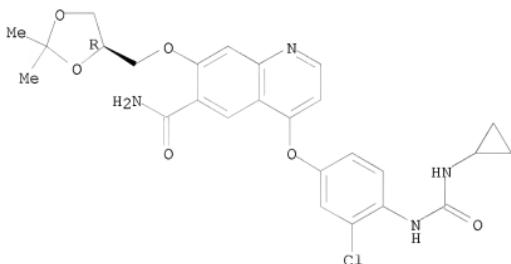
RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of urea derivs. containing nitrogenous aromatic ring compds. as angiogenesis inhibitors for prevention or treatment of diseases)

RN 417717-12-5 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(4R)-2,2-dimethyl-1,3-dioxolan-4-yl]methoxy]- (CA INDEX NAME)

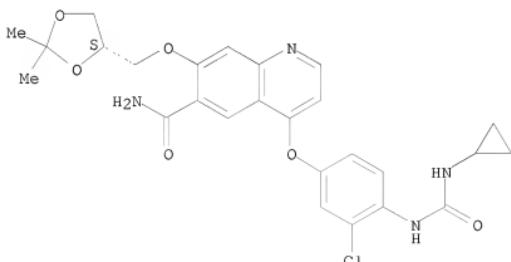
Absolute stereochemistry.



RN 417717-13-6 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(4S)-2,2-dimethyl-1,3-dioxolan-4-yl]methoxy]- (CA INDEX NAME)

Absolute stereochemistry.



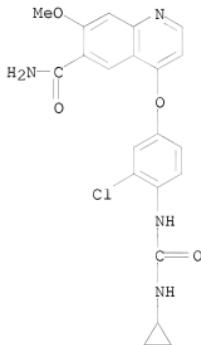
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 417717-14-7P 417717-15-8P 417717-16-9P
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 417719-50-7P 417719-56-3P 417719-57-4P
 417719-77-8P 417720-06-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of urea derivs. containing nitrogenous aromatic ring compds. as angiogenesis inhibitors for prevention or treatment of diseases)

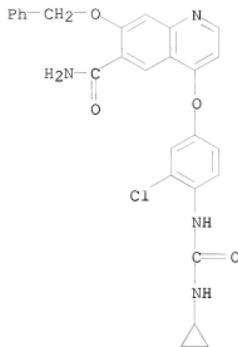
RN 417716-92-8 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)

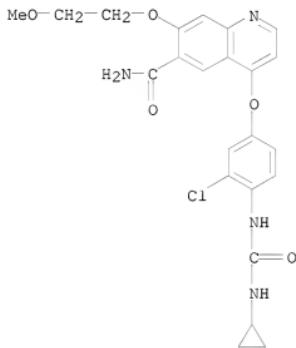


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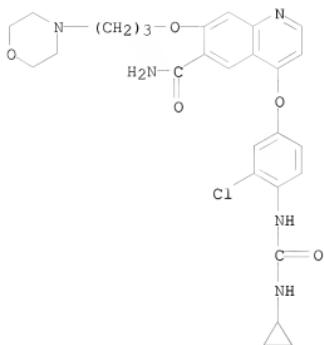
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(phenylmethoxy)- (CA INDEX NAME)



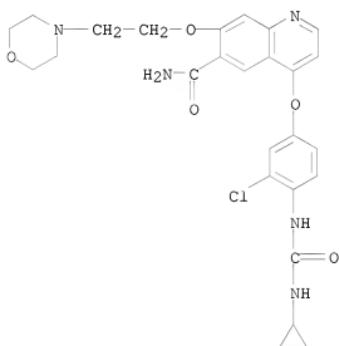
RN 417717-05-6 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



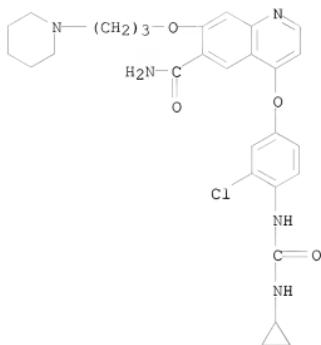
RN 417717-06-7 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[3-(4-morpholinyl)propoxy]- (CA INDEX NAME)



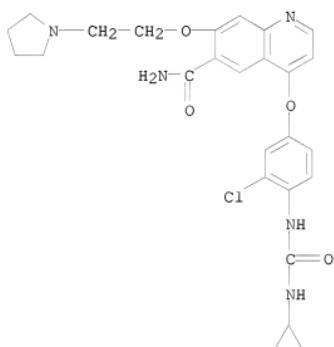
RN 417717-07-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)



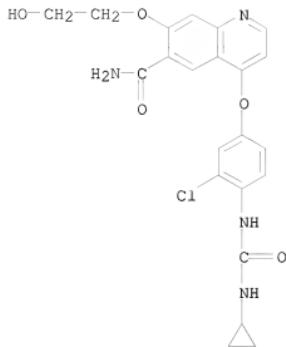
RN 417717-08-9 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[3-(1-piperidinyl)propoxy]- (CA INDEX NAME)



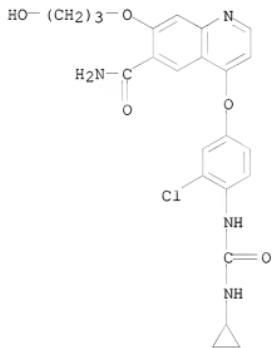
RN 417717-09-0 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(1-pyrrolidinyl)ethoxy]- (CA INDEX NAME)



RN 417717-10-3 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)

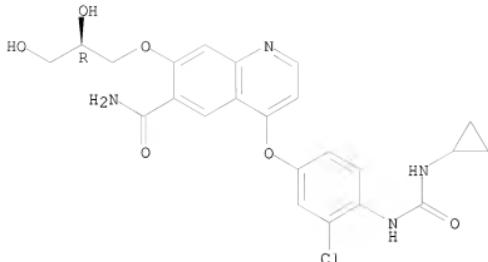


RN 417717-11-4 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(3-hydroxypropoxy)- (CA INDEX NAME)



RN 417717-14-7 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2,3-dihydroxypropoxy)- (CA INDEX NAME)

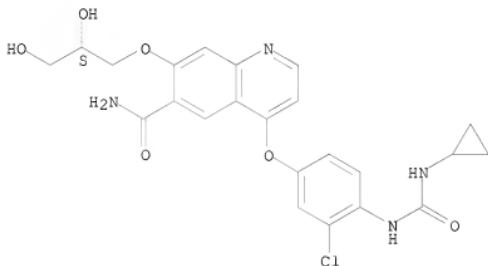
Absolute stereochemistry.



RN 417717-15-8 CA

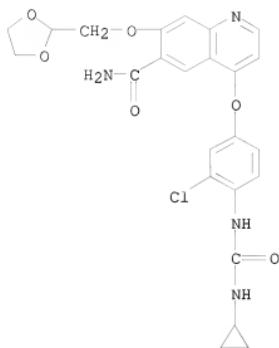
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.

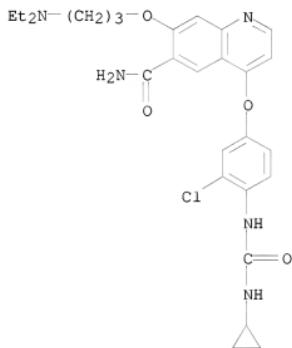


RN 417717-16-9 CA

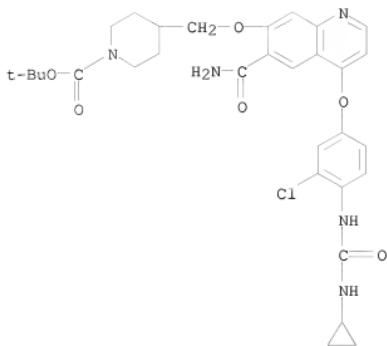
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(1,3-dioxolan-2-ylmethoxy)- (CA INDEX NAME)



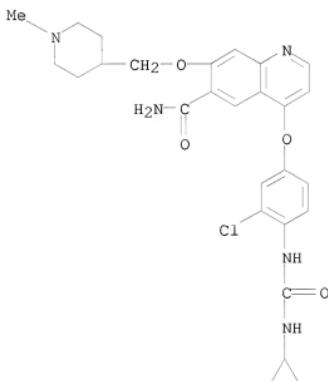
RN 417717-17-0 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[3-(diethylamino)propoxy]- (CA INDEX NAME)



RN 417717-18-1 CA
CN 1-Piperidinecarboxylic acid, 4-[[[6-(aminocarbonyl)-4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-quinolinyl]oxy]methyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

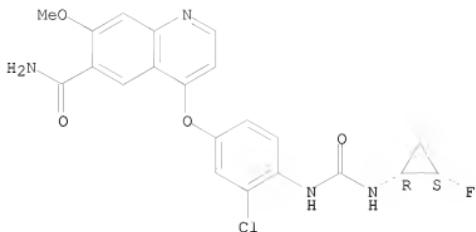


RN 417717-19-2 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy-7-[(1-methyl-4-piperidinyl)methoxy]- (CA INDEX NAME)



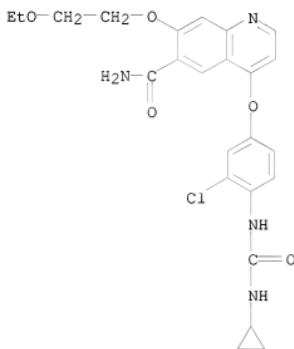
RN 417719-50-7 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropylamino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.



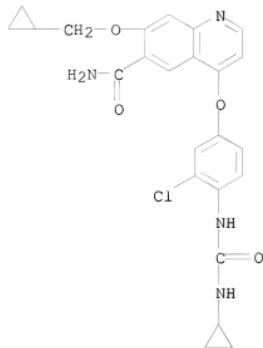
RN 417719-56-3 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



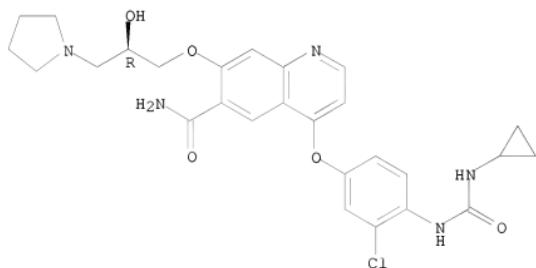
RN 417719-57-4 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(cyclopropylmethoxy)- (CA INDEX NAME)



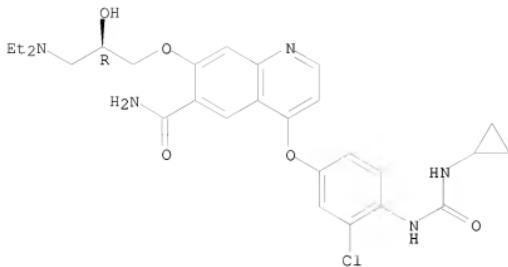
RN 417719-77-8 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

Absolute stereochemistry.



RN 417720-06-0 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-3-(diethylamino)-2-hydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.

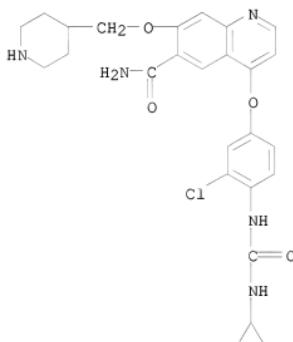


IT 417724-98-2P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of urea derivs. containing nitrogenous aromatic ring compds. as angiogenesis inhibitors for prevention or treatment of diseases)

RN 417724-98-2 CA

CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
heenoxy]-7-(4-piperidinylmethoxy)- (CA INDEX NAME)

REFERENCE COUNT:

117 THERE ARE 117 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 8 OF 22 CA COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 147:93969 CA

TITLE: Combination of anti-angiopoietin 2 human monoclonal antibody and of VEGF-A, KDR and/or FLT1 antagonist for treating cancer

INVENTOR(S): Brown, Jeffrey Lester; Emery, Stephen Charles; Blakey,
David Charles
 PATENT ASSIGNEE(S): Astrazeneca AB, Swed.; Astrazeneca UK Limited
 SOURCE: PCT Int. Appl., 88pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|------------|
| WO 2007068895 | A1 | 20070621 | WO 2006-GB4611 | 20061212 |
| WO 2007068895 | A9 | 20080612 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AP, EA, EP, OA | | | | |
| AU 2006324477 | A1 | 20070621 | AU 2006-324477 | 20061212 |
| EP 1962903 | A1 | 20080903 | EP 2006-820476 | 20061212 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, HR | | | | |
| KR 2008073766 | A | 20080811 | KR 2008-715745 | 20080627 |
| PRIORITY APPLN. INFO.: | | | US 2005-750551P | P 20051215 |
| | | | WO 2006-GB4611 | W 20061212 |

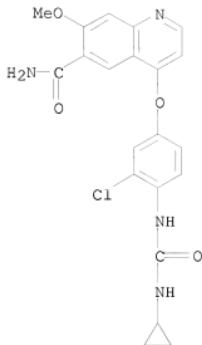
AB The invention relates to agents which possess anti-angiogenic activity and are accordingly useful in methods of treatment of disease states associated with angiogenesis in the animal or human body. More specifically the invention concerns a combination of a monoclonal antibody against human angiopoietin 2 (anti-Ang-2) and an antagonist of the biol. activity of VEGF-A, and/or KDR receptor, and/or FLT1, and uses of such antagonists. The nucleotide sequences and the encoded amino acid sequences of anti-Ang-2 monoclonal antibodies are disclosed.

IT 417716-92-8

RL: PAC (Pharmacological activity); BIOL (Biological study)
(combination of anti-angiopoietin 2 human monoclonal antibody and of VEGF-A, KDR and/or FLT1 antagonist for treating cancer)

RN 417716-92-8 CA

CN 6-Quinolincarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)

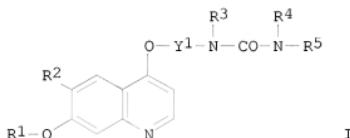


REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

| L4 ANSWER 9 OF 22 | CA | COPYRIGHT 2008 ACS on STN 147:23734 CA | | |
|---|------|---|------------------|------------|
| ACCESSION NUMBER: | | | | |
| TITLE: | | Anti-tumor agent for multiple myeloma | | |
| INVENTOR(S): | | Kamata, Junichi | | |
| PATENT ASSIGNEE(S): | | Eisai R & D Management Co., Ltd., Japan | | |
| SOURCE: | | PCT Int. Appl., 138pp. | | |
| | | CODEN: PIXXD2 | | |
| DOCUMENT TYPE: | | Patent | | |
| LANGUAGE: | | Japanese | | |
| FAMILY ACC. NUM. COUNT: | 2 | | | |
| PATENT INFORMATION: | | | | |
| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
| WO 2007061127 | A1 | 20070531 | WO 2006-JP323878 | 20061122 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| EP 1964837 | A1 | 20080903 | EP 2006-833681 | 20061122 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, RS | | | | |
| PRIORITY APPLN. INFO.: | | | JP 2005-337772 | A 20051122 |

US 2006-803450P P 20060530
 WO 2006-JP323878 W 20061122

OTHER SOURCE(S) : MARPAT 147:23734
 GI



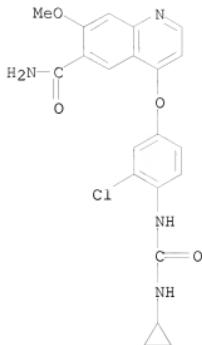
AB Disclosed is a pharmaceutical composition which can exert its effect with higher efficiency on a living body having at least one cell selected from the group consisting of a cell that over-expresses FGFR3, a cell that has a t(4;14) translocation and a cell that expresses a mutant FGFR3. Also disclosed is a therapeutic method for the living body. A compound represented by the general formula (I) or a pharmaceutically acceptable salt thereof or a solvate of the compound or the salt can exert its effect with higher efficiency on a living body having at least one cell selected from the group consisting of a cell that over-expresses FGFR3, a cell that has a t(4;14) translocation and a cell that expresses a mutant FGFR3.

IT
 417716-92-8 417717-05-6 417717-07-8
 417717-10-3 417717-15-8 417719-50-7
 417719-56-3 417719-77-8

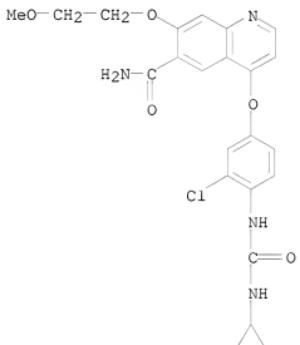
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (quinolin carboxamide analogs as FGFR3 inhibitors and antitumor agents for multiple myeloma)

RN 417716-92-8 CA

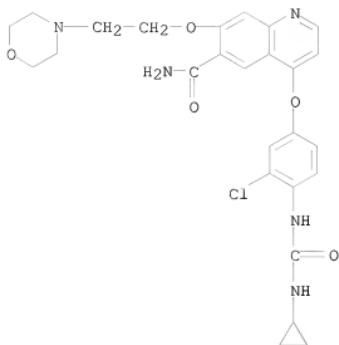
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



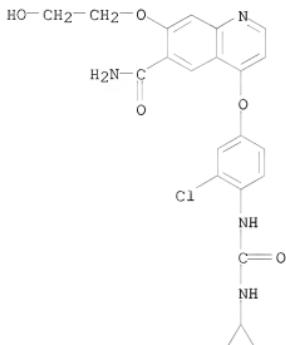
RN 417717-05-6 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-07-8 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

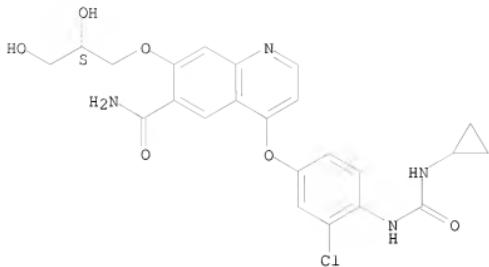


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamides, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



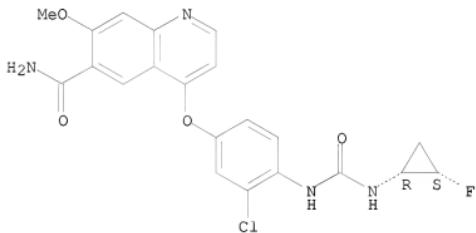
RN 417717-15-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy)- (CA INDEX NAME)

Absolute stereochemistry.

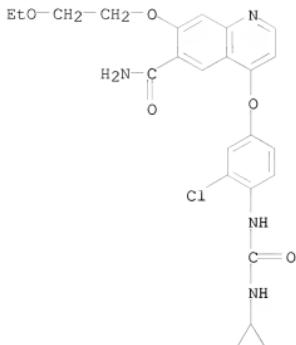


RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.

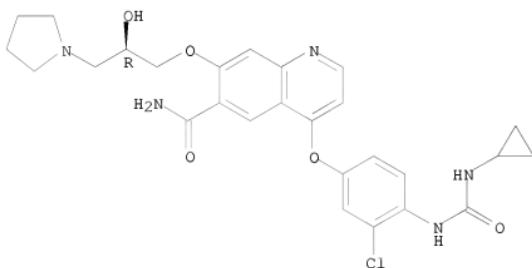


RN 417719-56-3 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-
 henoxy]-7-[{(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

Absolute stereochemistry.



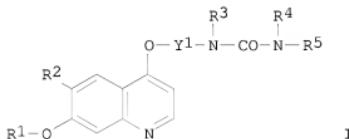
REFERENCE COUNT: 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 10 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 147:23732 CA
 TITLE: Anti-tumor agent for multiple myeloma
 INVENTOR(S): Kamata, Junichi
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 139pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------------------------|--------------------------|
| WO 2007061130 | A1 | 20070531 | WO 2006-JP323881 | 20061122 |
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| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| PRIORITY APPLN. INFO.: | | | JP 2005-337772 US 2006-803450P | A 20051122 P 20060530 |

OTHER SOURCE(S): MARPAT 147:23732
GI

AB Disclosed is a pharmaceutical composition which can exert its effect with higher efficiency on a living body having at least one cell selected from the group consisting of a cell that over-expresses FGFR3, a cell that has a t(4;14) translocation and a cell that expresses a mutant FGFR3. Also disclosed is a therapeutic method for the living body. A compound represented by the general formula (I) or a pharmaceutically acceptable salt thereof or a solvate of the compound or the salt can exert its effect with higher efficiency on a living body having at least one cell selected from the group consisting of a cell that over-expresses FGFR3, a cell that has a t(4;14) translocation and a cell that expresses a mutant FGFR3.

IT 417716-92-8 417717-05-6 417717-07-8

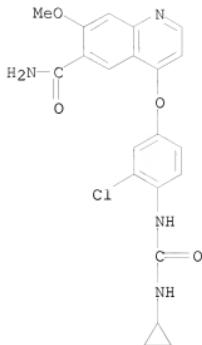
417717-10-3 417717-15-8 417719-50-7

417719-56-3 417719-77-8

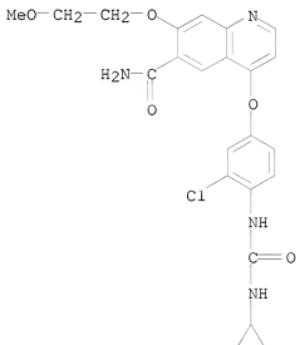
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(quinolin carboxamide analogs as FGFR3 inhibitors and antitumor agents for multiple myeloma)

RN 417716-92-8 CA

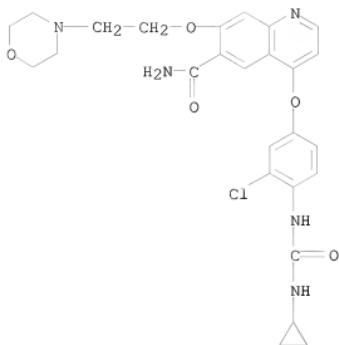
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy-7-methoxy- (CA INDEX NAME)



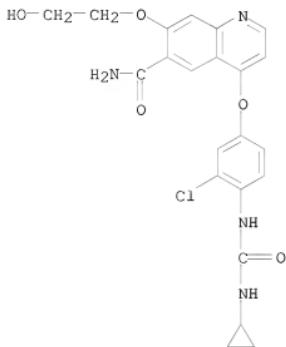
RN 417717-05-6 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-07-8 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

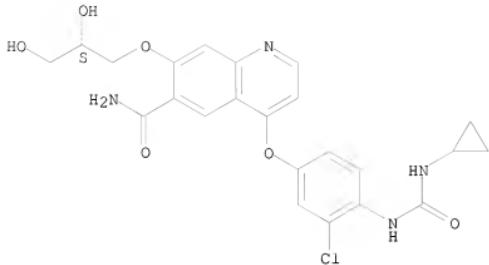


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamides, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



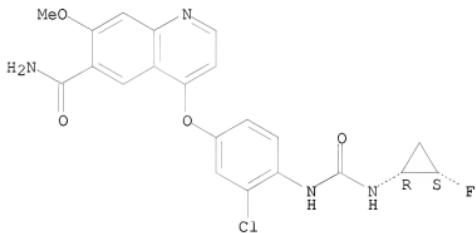
RN 417717-15-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.

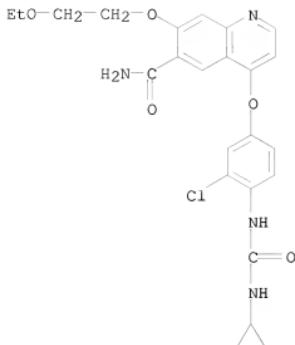


RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.

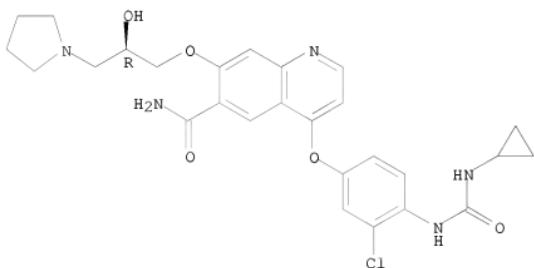


RN 417719-56-3 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-henoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 11 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 146:455231 CA
 TITLE: Use of combination of anti-angiogenic substance and c-kit kinase inhibitor
 INVENTOR(S): Yamamoto, Yuji
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 102pp.
 CODEN: PIXD2
 DOCUMENT TYPE: Patent

LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------------------------|--------------------------|
| WO 2007052850 | A1 | 20070510 | WO 2006-JP322516 | 20061107 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| EP 1949902 | A1 | 20080730 | EP 2006-832529 | 20061107 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, RS | | | | |
| PRIORITY APPLN. INFO.: | | | JP 2005-322946 WO 2006-JP322516 | A 20051107 W 20061107 |

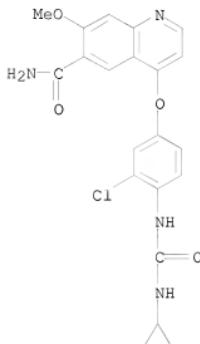
OTHER SOURCE(S): MARPAT 146:455231

AB Disclosed are a pharmaceutical composition having an excellent anti-tumor effect, and a therapeutic method for cancer. 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7'-methoxy-6-quinolinecarboxamide or an analog thereof can be used in combination with a substance having a c-kit kinase-inhibiting activity to produce an excellent anti-tumor effect. For example, the effect of combination of 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7'-methoxy-6-quinolinecarboxamide methanesulfonate and imatinib on human gastrointestinal stromal tumor cell (GIST882 cell)-bearing model mice was examined

IT 417716-92-8, 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-methoxy-6-quinolinecarboxamide 417717-05-6,
 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-(2-methoxyethoxy)-6-quinolinecarboxamide 417717-07-8, 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-[2-(4-morpholino)ethoxy]-6-quinolinecarboxamide 417717-10-3, 4-(3-Chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)-6-quinolinecarboxamide 417717-15-8, 4-(3-Chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy)-7-(2S)-2,3-dihydroxypropyl oxy-6-quinolinecarboxamide 417719-50-7,
 4-[3-Chloro-4-(cis-2-fluoro-cyclopropylaminocarbonyl)aminophenoxy]-7-methoxy-6-quinolinecarboxamide 417719-56-3, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-ethoxyethoxy)-6-quinolinecarboxamide 417719-77-8, 4-[3-Chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2R)-2-hydroxy-3-(1-pyrrolidino)propoxy]-6-quinolinecarboxamide 857890-39-2
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (use of combination of anti-angiogenic substance and c-kit kinase inhibitor)

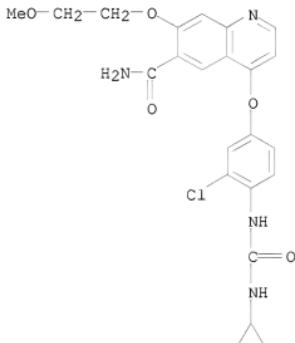
RN 417716-92-8 CA
 CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]

heoxy]-7-methoxy- (CA INDEX NAME)



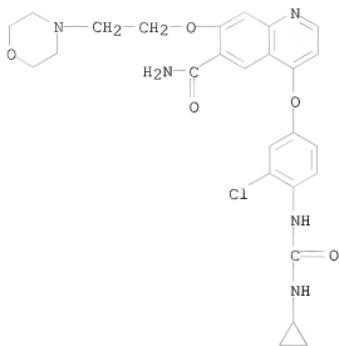
RN 417717-05-6 CA

CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]heoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)

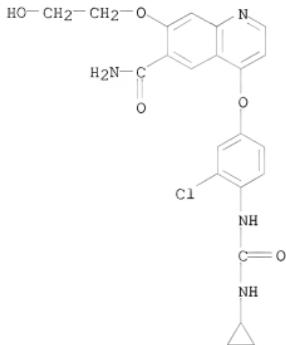


RN 417717-07-8 CA

CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]heoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

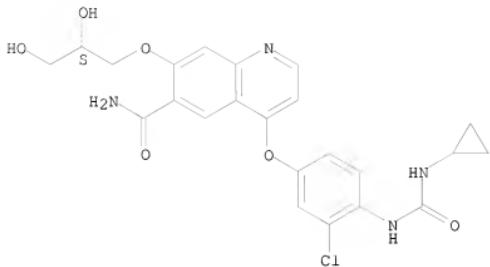


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



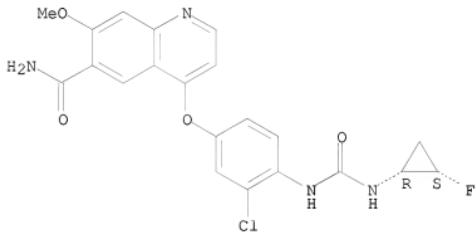
RN 417717-15-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy)- (CA INDEX NAME)

Absolute stereochemistry.

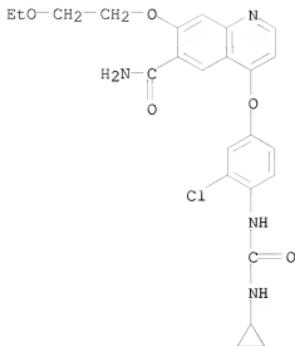


RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.

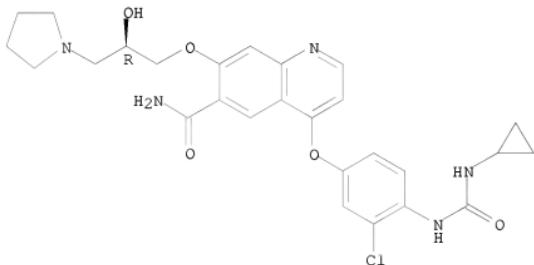


RN 417719-56-3 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

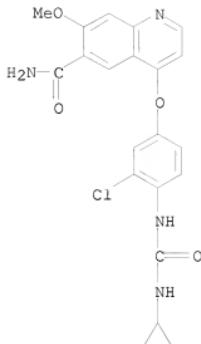
Absolute stereochemistry.



RN 857890-39-2 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
 CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2
CMF C H4 O3 S

REFERENCE COUNT: 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 12 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 146:455230 CA
 TITLE: Use of combination of anti-angiogenic substance and c-kit kinase inhibitor
 Yamamoto, Yuji
 INVENTOR(S):
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 103pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|------------------|----------|
| WO 2007052849 | A1 | 20070510 | WO 2006-JP322514 | 20061107 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, | | | | |

KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK,
 MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO,
 RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT,
 TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW

RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
 IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ,
 CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
 GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
 KG, KZ, MD, RU, TJ, TM

AU 2006309551

A1 20070510

AU 2006-309551

20061107

CA 2627598

A1 20070510

CA 2006-2627598

20061107

KR 2008065698

A 20080714

KR 2008-713685

20080605

PRIORITY APPLN. INFO.:

JP 2005-322946

A 20051107

WO 2006-JP322514

W 20061107

OTHER SOURCE(S): MARPAT 146:455230

AB Disclosed are a pharmaceutical composition having an excellent anti-tumor effect, and a therapeutic method for cancer. 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7'-methoxy-6-quinolinicarboxamide or an analog thereof can be used in combination with a substance having a c-kit kinase-inhibiting activity to produce an excellent anti-tumor effect. For example, the effect of combination of 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7'-methoxy-6-quinolinicarboxamide methanesulfonate and imatinib on human gastrointestinal stromal tumor cell (GIST882 cell)-bearing model mice was examined

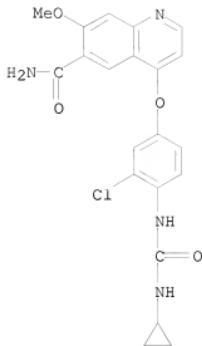
IT 417716-92-8, 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-methoxy-6-quinolinicarboxamide 417717-05-6,
 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-(2-methoxyethoxy)-6-quinolinicarboxamide 417717-07-8, 4-[3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy]-7-[2-(4-morpholino)ethoxy-6-quinolinicarboxamide 417717-10-3, 4-(3-Chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)-6-quinolinicarboxamide 417717-15-8, 4-(3-Chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2S]-2,3-dihydroxypropyl)oxy-6-quinolinicarboxamide 417719-50-7,
 4-[3-Chloro-4-(cis-2-fluoro-cyclopropylaminocarbonyl)aminophenoxy]-7-methoxy-6-quinolinicarboxamide 417719-56-3, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-ethoxyethoxy)-6-quinolinicarboxamide 417719-77-8, 4-[3-Chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidino)propoxy]-6-quinolinicarboxamide 857890-39-2

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

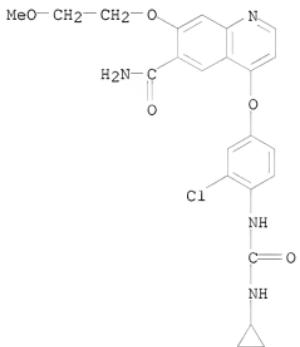
(use of combination of anti-angiogenic substance and c-kit kinase inhibitor)

RN 417716-92-8 CA

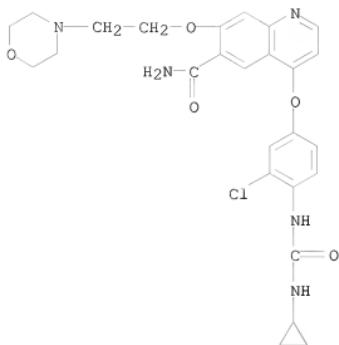
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



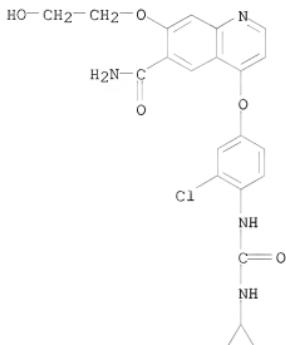
RN 417717-05-6 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-07-8 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

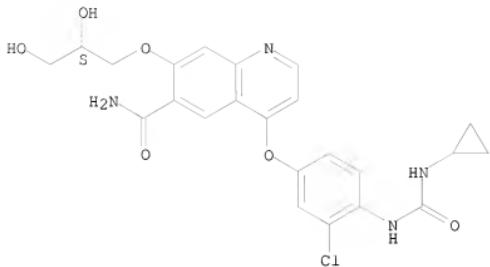


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamides, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



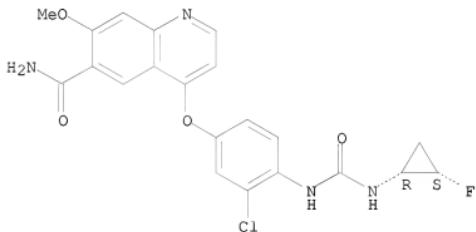
RN 417717-15-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.

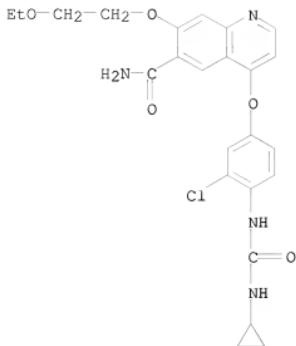


RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.

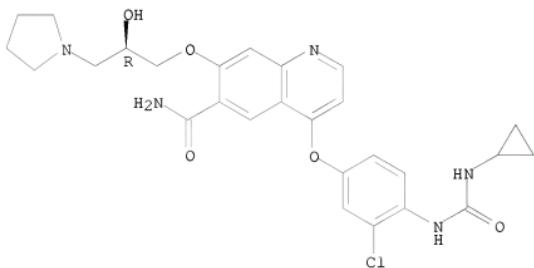


RN 417719-56-3 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

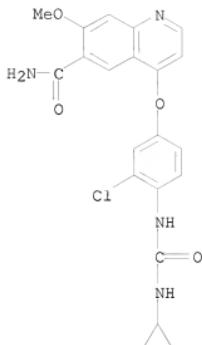
Absolute stereochemistry.



RN 857890-39-2 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
 CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2
CMF C H4 O3 S

REFERENCE COUNT: 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 13 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 146:221063 CA
 TITLE: Method for assaying anti-tumor effect of angiogenesis inhibitor
 INVENTOR(S): Uenaka, Toshimitsu; Yamamoto, Yuji; Matsui, Junji
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 147pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|------------------|----------|
| WO 2007015578 | A1 | 20070208 | WO 2006-JP315698 | 20060802 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, | | | | |

KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN,
 MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU,
 SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG,
 US, UZ, VC, VN, ZA, ZM, ZW

RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
 IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ,
 CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
 GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
 KG, KZ, MD, RU, TJ, TM

EP 1925676 A1 20080528 EP 2006-768437 20060802

R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
 IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL,
 BA, HR, MK, RS

PRIORITY APPLN. INFO.:

JP 2005-224173 A 20050802
 JP 2006-164700 A 20060514
 WO 2006-JP315698 W 20060802

OTHER SOURCE(S): MARPAT 146:221063

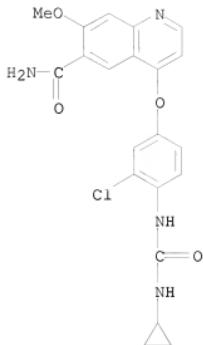
AB Disclosed is a method for predicting the anti-tumor effect of an angiogenesis inhibitor. The method comprises evaluating the EGF-dependence property of an angiogenesis inhibitor with respect to proliferation and/or survival of tumor cells, and using the evaluated EGF-dependence property as a measure. The anti-tumor effect of an angiogenesis inhibitor correlates with the EGF-dependency property of the inhibitor with respect to proliferation and/or survival of tumor cells. Therefore, an angiogenesis inhibitor is capable of exerting an excellent anti-tumor effect by using it in combination with a substance having an EGF inhibitory effect.

IT 417716-92-8, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide 417716-92-8D,
 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide, pharmacol. allowed salt, solvate 417717-05-6
 , 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-methoxyethoxy)-6-quinolinecarboxamide 417717-07-8
 417717-10-3 417717-15-8 417719-50-7
 417719-56-3 417719-77-8 857890-39-2

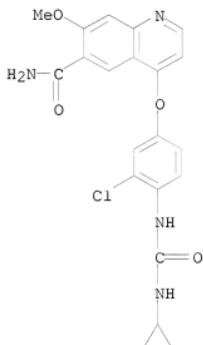
RL: ANT (Analyte); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
 (method for assaying anti-tumor effect of angiogenesis inhibitor by evaluating EGF-dependency)

RN 417716-92-8 CA

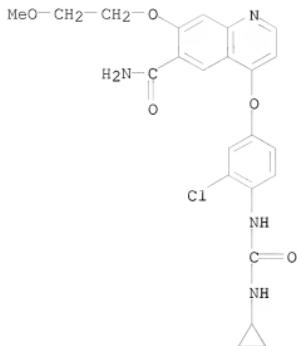
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



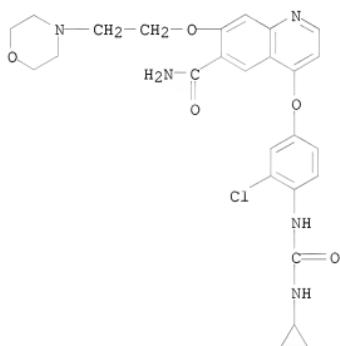
RN 417716-92-8 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-methoxy- (CA INDEX NAME)



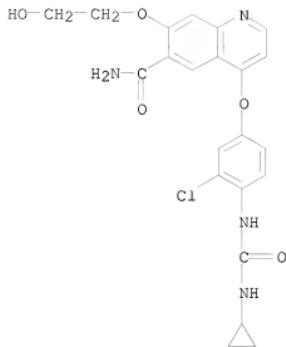
RN 417717-05-6 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-07-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

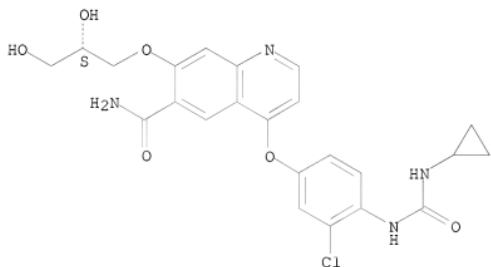


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



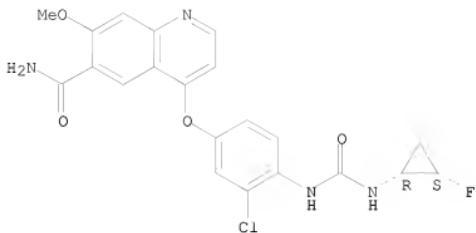
RN 417717-15-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.

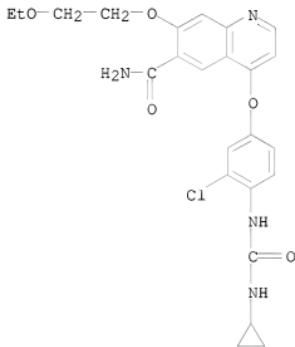


RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.

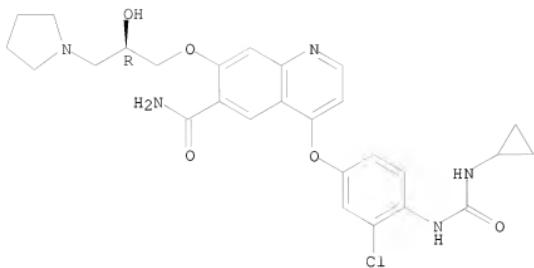


RN 417719-56-3 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

Absolute stereochemistry.



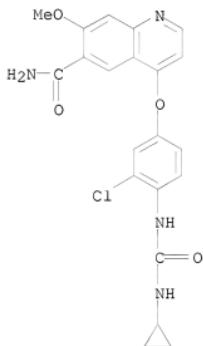
RN 857890-39-2 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8

CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2

CMF C H4 O3 S



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 14 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 146:221062 CA
 TITLE: Method for predicting antitumor efficacy of angiogenesis inhibitor
 INVENTOR(S): Matsui, Junji; Semba, Taro
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 104pp.
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|------------|
| WO 2007015569 | A1 | 20070208 | WO 2006-JP315563 | 20060801 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| EP 1925941 | A1 | 20080528 | EP 2006-782407 | 20060801 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, RS | | | | |
| PRIORITY APPLN. INFO.: | | | JP 2005-223440 | A 20050801 |
| | | | WO 2006-JP315563 | W 20060801 |

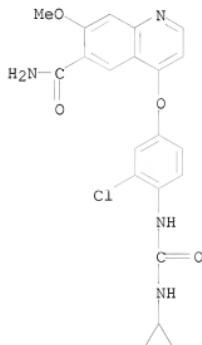
OTHER SOURCE(S): MARPAT 146:221062
 AB A method for predicting the antitumor efficacy of an angiogenesis inhibitor is provided, which comprises measuring the number of blood vessels surrounded by pericytes in tumor, and using the measurement value as a measure for the anti-tumor effect.
 IT 417716-92-8, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide 417716-92-8D,
 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide, pharmacol. allowed salt, solvate 417717-05-6
 , 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-methoxyethoxy)-6-quinolinecarboxamide 417717-07-8
 417717-10-3 417717-15-8 417719-50-7

417719-56-3 417719-77-8 857890-39-2

RL: ANT (Analyte); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
 (method for predicting antitumor efficacy of angiogenesis inhibitor)

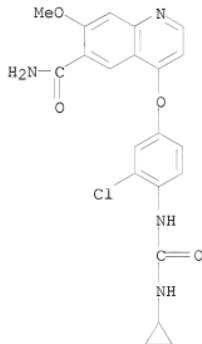
RN 417716-92-8 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
 henoxy-7-methoxy- (CA INDEX NAME)



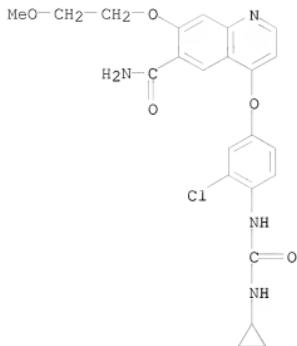
RN 417716-92-8 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
 henoxy-7-methoxy- (CA INDEX NAME)

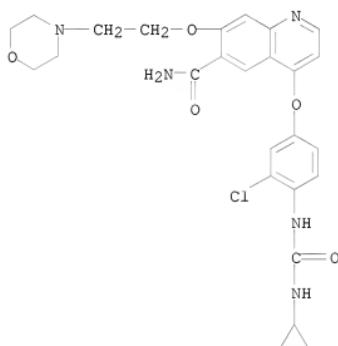


RN 417717-05-6 CA

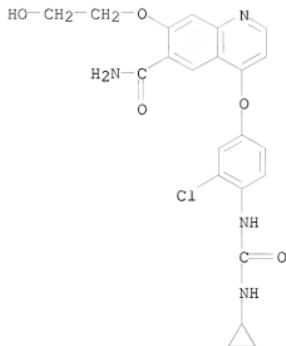
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
 henoxy-7-(2-methoxyethoxy)- (CA INDEX NAME)



RN 417717-07-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

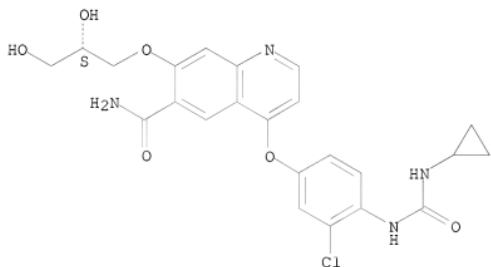


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



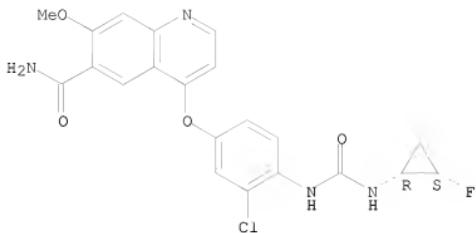
RN 417717-15-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



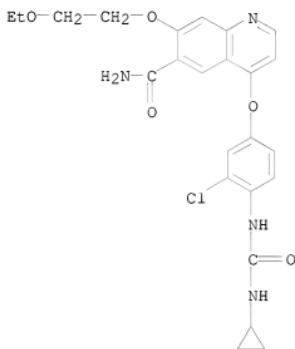
RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.



RN 417719-56-3 CA

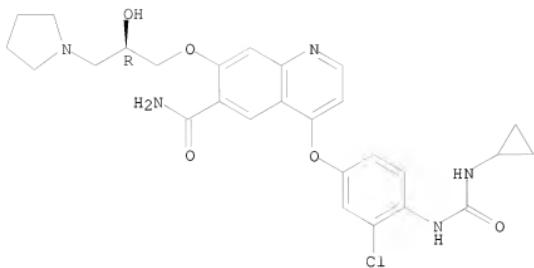
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA

CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[{(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy}-] (CA INDEX NAME)

Absolute stereochemistry.



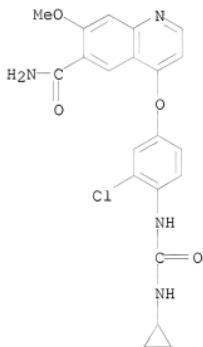
RN 857890-39-2 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]oxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8

CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2

CMF C H4 O3 S

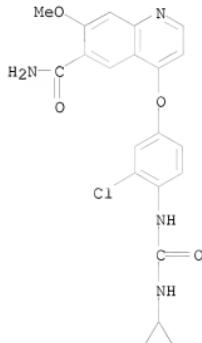


REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

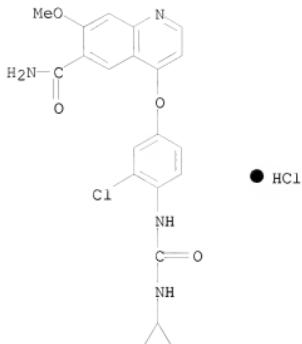
L4 ANSWER 15 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 146:100576 CA
 TITLE: Preparation of amorphous salts of 4-[3-chloro-4-[(cyclopropylaminocarbonyl)amino]phenoxy]-7-methoxy-6-quinolinecarboxamide as antitumor agents
 INVENTOR(S): Sakaguchi, Takahisa; Tsuruoka, Akihiko
 PATENT ASSIGNEE(S): Eisai R & D Management Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 49pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|------------|
| WO 2006137474 | A1 | 20061228 | WO 2006-JP312487 | 20060622 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| AU 2006260148 | A1 | 20061228 | AU 2006-260148 | 20060622 |
| CA 2606719 | A1 | 20061228 | CA 2006-2606719 | 20060622 |
| US 20070004773 | A1 | 20070104 | US 2006-472372 | 20060622 |
| EP 1894918 | A1 | 20080305 | EP 2006-767145 | 20060622 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, YU | | | | |
| KR 2008008374 | A | 20080123 | KR 2007-727079 | 20071121 |
| CN 101233111 | A | 20080730 | CN 2006-80020317 | 20071207 |
| PRIORITY APPLN. INFO.: | | | US 2005-693044P | P 20050623 |
| | | | WO 2006-JP312487 | W 20060622 |
| AB This invention pertains to a method for producing amorphous salts of 4-[3-chloro-4-[(cyclopropylaminocarbonyl)amino]phenoxy]-7-methoxy-6-quinolinecarboxamide. The title compds. are useful as antitumor agents for various cancers, such as pancreas cancer, stomach cancer, colon cancer, breast cancer, prostate cancer, lung cancer, renal cancer, brain cancer, blood cancer, ovarian cancer, and hemangioma (no data). | | | | |

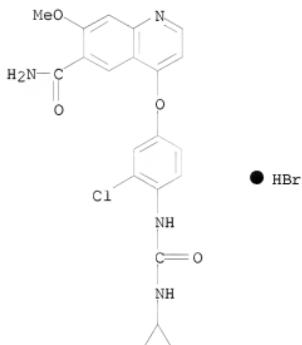
IT 417716-92-8P
 RL: PAC (Pharmacological activity); PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (drug candidate; preparation of salts of 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-6-quinolinecarboxamide as antitumor agents)
 RN 417716-92-8 CA
 CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



IT 857890-31-4P 857890-33-6P 857890-35-8P
 857890-37-0P 857890-39-2P 857890-41-6P
 857890-45-0P 857890-47-2P 917572-43-1P
 917572-44-2P
 RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (drug candidate; preparation of salts of 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-6-quinolinecarboxamide as antitumor agents)
 RN 857890-31-4 CA
 CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, hydrochloride (1:1) (CA INDEX NAME)



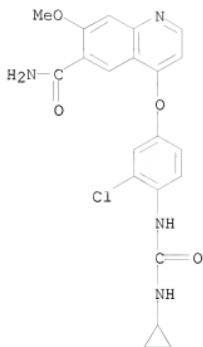
RN 857890-33-6 CA
 CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-henoxy]-7-methoxy-, hydrobromide (1:1) (CA INDEX NAME)



RN 857890-35-8 CA
 CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-henoxy]-7-methoxy-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

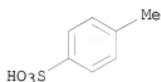
CM 1

CRN 417716-92-8
 CMF C21 H19 Cl N4 O4



CM 2

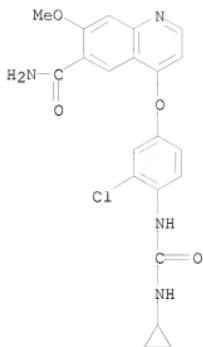
CRN 104-15-4
CMF C7 H8 O3 S



RN 857890-37-0 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, sulfate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

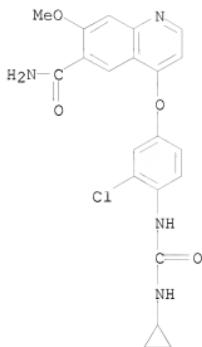
CRN 7664-93-9
CMF H₂ O₄ S



RN 857890-39-2 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxyl-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C₂₁ H₁₉ Cl N₄ O₄

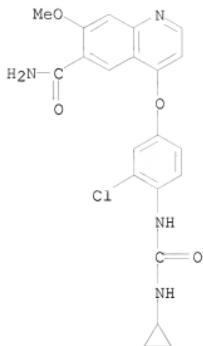


CM 2

CRN 75-75-2
CMF C H4 O3 SRN 857890-41-6 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]hexyloxy]-7-methoxy-, methanesulfonate, hydrate (1:1:?) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

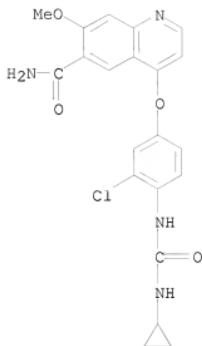
CRN 75-75-2
CMF C H4 O3 S



RN 857890-45-0 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]heoxy]-7-methoxy-, acetate compd. with methanesulfonic acid (1?:?) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2
CMF C H4 O3 S



CM 3

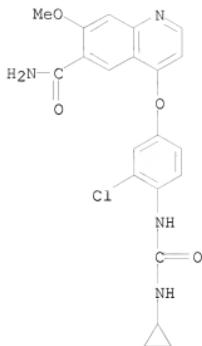
CRN 64-19-7
CMF C2 H4 O2



RN 857890-47-2 CA
CN Ethanesulfonic acid, compd. with 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-6-quinolinecarboxamide (1:1) (CA INDEX NAME)

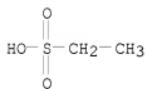
CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

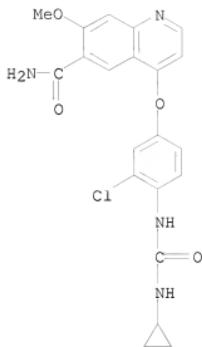
CRN 594-45-6
CMF C2 H6 O3 S



RN 917572-43-1 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]hexyloxy]-7-methoxy-, methanesulfonate, compd. with 1,1'-sulfinylbis[methane] (1:1:?) (CA INDEX NAME)

CM 1

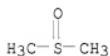
CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2
CMF C H4 O3 S

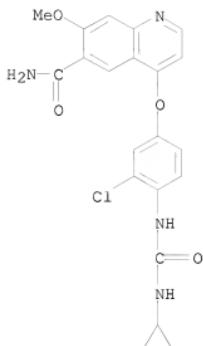
CM 3

CRN 67-68-5
CMF C2 H6 O S

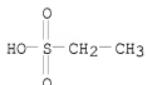
RN 917572-44-2 CA
 CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]heoxy]-7-methoxy-, ethanesulfonate, compd. with 1,1'-sulfinylbis[methane] (1:1:?) (CA INDEX NAME)

CM 1

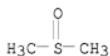
CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

CRN 594-45-6
CMF C2 H6 O3 S

CM 3

CRN 67-68-5
CMF C2 H6 O S

REFERENCE COUNT: 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

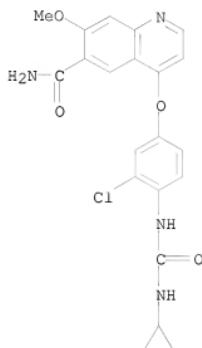
L4 ANSWER 16 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 144:324798 CA
 TITLE: Simultaneous use of sulfonamide-containing compound and angiogenesis inhibitor
 Owa, Takashi; Ozawa, Yoichi; Semba, Taro
 Eisai Co., Ltd., Japan
 INVENTOR(S):
 PATENT ASSIGNEE(S):
 SOURCE: PCT Int. Appl., 270 pp.
 CODEN: PIXXD2

DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 6
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|------------|
| WO 2006030941 | A1 | 20060323 | WO 2005-JP17228 | 20050913 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| WO 2006030947 | A1 | 20060323 | WO 2005-JP17238 | 20050913 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| US 20060135486 | A1 | 20060622 | US 2005-226655 | 20050913 |
| EP 1797877 | A1 | 20070620 | EP 2005-785820 | 20050913 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, YU | | | | |
| PRIORITY APPLN. INFO.: | | | US 2004-609452P | P 20040913 |
| | | | JP 2005-54150 | A 20050228 |
| | | | JP 2005-54475 | A 20050228 |
| | | | WO 2005-JP17238 | W 20050913 |

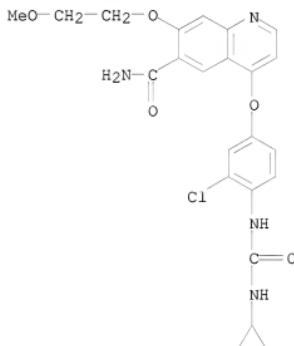
OTHER SOURCE(S): MARPAT 144:324798
 AB A pharmaceutical composition comprising a sulfonamide-containing compound combined with an angiogenesis inhibitor.
 IT 417716-92-8, 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinicarboxamide 417717-05-6,
 4-(3-Chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-(2-methoxyethoxy)-6-quinolinicarboxamide 417717-07-8 417717-10-3
 417717-15-8 417719-50-7 417719-56-3
 417719-77-8
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (sulfonamide-containing compds. and angiogenesis inhibitors for combination chemotherapy of cancer)
 RN 417716-92-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-methoxy- (CA INDEX NAME)



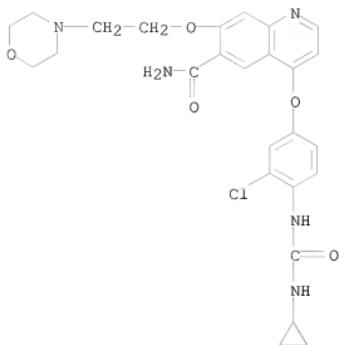
RN 417717-05-6 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)

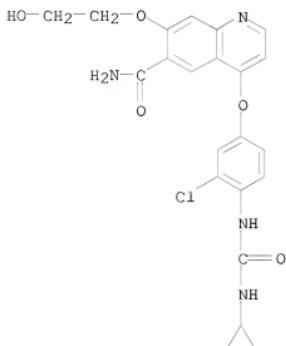


RN 417717-07-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)

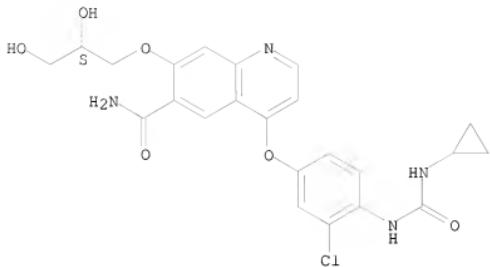


RN 417717-10-3 CA
CN 6-Quinolinemcarboxamides, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)



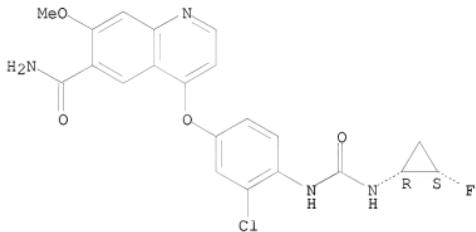
RN 417717-15-8 CA
CN 6-Quinolinemcarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy)- (CA INDEX NAME)

Absolute stereochemistry.

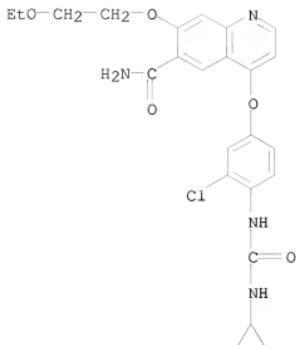


RN 417719-50-7 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy]-7-methoxy-, rel- (CA INDEX NAME)

Relative stereochemistry.

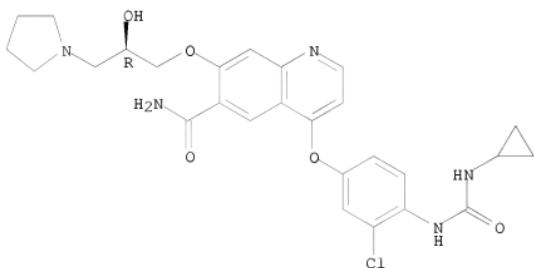


RN 417719-56-3 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]pro-
 henoxy]-7-[(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy]- (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 17 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 144:299488 CA
 TITLE: Stable medicinal compositions of quinoliniccarboxamide derivative
 INVENTOR(S): Furitsu, Hisao; Suzuki, Yasuyuki
 PATENT ASSIGNEE(S): Eisai Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 53 pp.
 CODEN: PIXDZ
 DOCUMENT TYPE: Patent

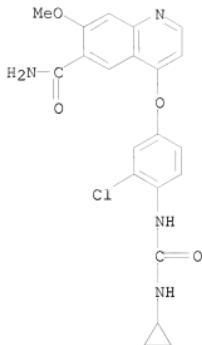
LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|------------|
| WO 2006030826 | A1 | 20060323 | WO 2005-JP16941 | 20050914 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MM, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| AU 2005283422 | A1 | 20060323 | AU 2005-283422 | 20050914 |
| CA 2579810 | A1 | 20060323 | CA 2005-2579810 | 20050914 |
| EP 1797881 | A1 | 20070620 | EP 2005-783232 | 20050914 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, YU | | | | |
| CN 101001629 | A | 20070718 | CN 2005-80026468 | 20050914 |
| KR 2007053205 | A | 20070523 | KR 2007-701347 | 20070119 |
| IN 2007CN01571 | A | 20070831 | IN 2007-CN1571 | 20070417 |
| US 20080214604 | A1 | 20080904 | US 2008-662425 | 20080404 |
| PRIORITY APPLN. INFO.: | | | JP 2004-272625 | A 20040917 |
| | | | WO 2005-JP16941 | W 20050914 |

AB This invention relates to highly stable medicinal composition which comprises 4-(3-chloro-4-(cyclopropylamino-carbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide (I), salts or solvates thereof, a compound whose 5 % aqueous solution or dispersion has a pH of 8 or higher, and/or silicic acid, salts or solvates thereof. Decomposition and surface gelation of I during storage at high humidity and temperature, is prevented. For example, tablets were formulated containing I-methanesulfonate salt 24, Aerosil-200 192, mannitol 1236, Avicel PH101 720, hydroxypropyl cellulose 72, Ac-Di-Sol 120, Na stearyl fumarate 36 parts and coated with Opadry Yellow.

IT 417716-92-8P, 4-(3-Chloro-4-(cyclopropylamino-carbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of quinolinecarboxamide derivative)

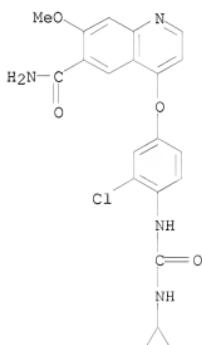
RN 417716-92-8 CA
 CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



IT 857890-39-2
RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES
(Uses)
(preparation of quinolinecarboxamide derivative and stable tablets
containing the
same)
RN 857890-39-2 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxo-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4

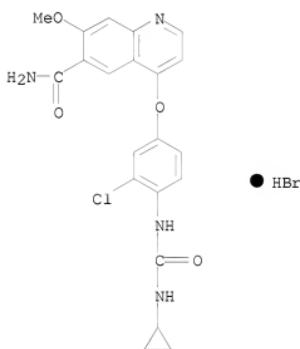


CM 2

CRN 75-75-2
CMF C H4 O3 SIT 857890-33-6 857890-35-8 857890-37-0
857890-41-6 857890-43-8 857890-45-0
857890-47-2RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(preparation of quinolinecarboxamide derivative and stable tablets containing the same)

RN 857890-33-6 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, hydrobromide (1:1) (CA INDEX NAME)

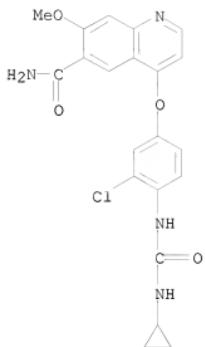


RN 857890-35-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

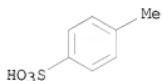
CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

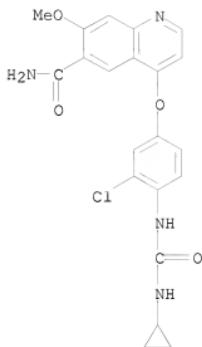
CRN 104-15-4
CMF C7 H8 O3 S



RN 857890-37-0 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, sulfate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

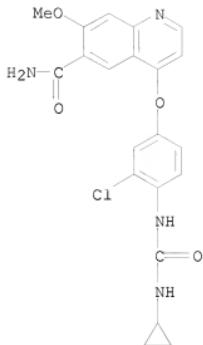
CRN 7664-93-9
CMF H₂ O₄ S



RN 857890-41-6 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-henoxy]-7-methoxy-, methanesulfonate, hydrate (1:1:?) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C₂₁ H₁₉ Cl N₄ O₄



CM 2

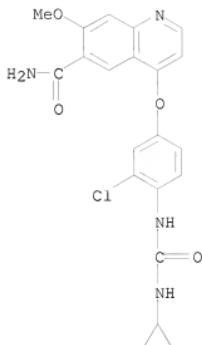
CRN 75-75-2
CMF C H4 O3 S



RN 857890-43-8 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-henoxy]-7-methoxy-, monomethanesulfonate, compd. with sulfonylbis[methane] (9CI) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



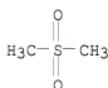
CM 2

CRN 75-75-2
 CMF C H4 O3 S



CM 3

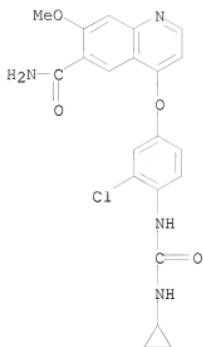
CRN 67-71-0
 CMF C2 H6 O2 S



RN 857890-45-0 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]heoxy-7-methoxy-, acetate compd. with methanesulfonic acid (1:?:?) (CA INDEX NAME)

CM 1

CRN 417716-92-8
 CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2
CMF C H4 O3 S



CM 3

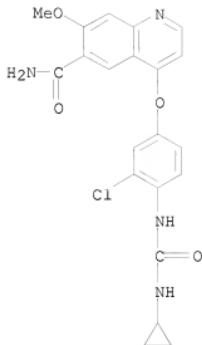
CRN 64-19-7
CMF C2 H4 O2



RN 857890-47-2 CA
CN Ethanesulfonic acid, compd. with 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-6-quinolinecarboxamide (1:1) (CA INDEX NAME)

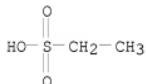
CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

CRN 594-45-6
CMF C2 H6 O3 S

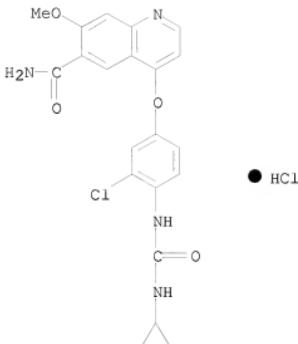


IT 857890-31-4

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(stable tablets containing quinolinecarboxamide derivative and alkalies and silicates)

RN 857890-31-4 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-henoxy]-7-methoxy-, hydrochloride (1:1) (CA INDEX NAME)

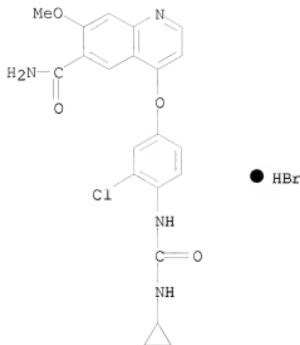


REFERENCE COUNT: 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 18 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 143:120562 CA
 TITLE: Crystal of salt of 4-[3-chloro-4-(cyclopropylaminocarbonyl)amino-phenoxy]-7-methoxy-6-quinolinicarboxamide or solvate thereof and processes for producing these
 INVENTOR(S): Matsushima, Tomohiro; Nakamura, Taiju; Yoshizawa, Kazuhiro; Kamada, Atsushi; Ayata, Yusuke; Suzuki, Naoko; Arimoto, Itaru; Sakaguchi, Takahisa; Gotoda, Masaharu
 PATENT ASSIGNEE(S): Eisai Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 95 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2005063713 | A1 | 20050714 | WO 2004-JP19223 | 20041222 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |

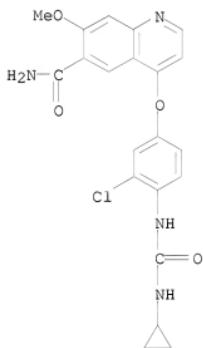
| | | | | |
|---|--|----------|------------------|-------------|
| AU 2004309217 | A1 | 20050714 | AU 2004-309217 | 20041222 |
| CA 2543650 | A1 | 20050714 | CA 2004-2543650 | 20041222 |
| EP 1698623 | A1 | 20060906 | EP 2004-807580 | 20041222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, BA, HR, IS, YU | | | | |
| CN 1890220 | A | 20070103 | CN 2004-80036184 | 20041222 |
| BR 2004018200 | A | 20070417 | BR 2004-18200 | 20041222 |
| RU 2328489 | C2 | 20080710 | RU 2006-126977 | 20041222 |
| US 20070078159 | A1 | 20070405 | US 2006-577531 | 20060428 |
| MX 2006PA07256 | A | 20060823 | MX 2006-PA7256 | 20060622 |
| KR 804566 | B1 | 20080220 | KR 2006-713993 | 20060712 |
| IN 2006CN02572 | A | 20070608 | IN 2006-CN2572 | 20060713 |
| NO 2006003383 | A | 20060925 | NO 2006-3383 | 20060721 |
| KR 2007107185 | A | 20071106 | KR 2007-722490 | 20071001 |
| KR 2008028511 | A | 20080331 | KR 2008-705282 | 20080303 |
| KR 839554 | B1 | 20080620 | | |
| PRIORITY APPLN. INFO.: | | | JP 2003-430939 | A 20031225 |
| | | | WO 2004-JP19223 | W 20041222 |
| | | | KR 2006-713993 | A3 20060712 |
| | | | KR 2007-722490 | A3 20071001 |
| AB | Disclosed are crystals of the hydrochloride, hydrobromide, p-toluenesulfonate, sulfate, methanesulfonate, or ethanesulfonate of 4-[3-chloro-4-(cyclopropylamino-carbonyl)aminophenoxy]-7-methoxy-6-quinolinecarboxamide or crystals of a solvate of any of these. The crystals have improved physicochem. and pharmacokinetic properties, and suitable for use as neovascularization inhibitors for treatment of related diseases. | | | |
| IT | 857890-33-6P 857890-39-2P RL: PEP (Physical, engineering or chemical process); PKT (Pharmacokinetics); PRP (Properties); PYP (Physical process); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (crystal of salt of 4-[3-chloro-4-(cyclopropylaminocarbonyl)amino-phenoxy]-7-methoxy-6-quinolinecarboxamide or solvate thereof as neovascularization inhibitor, and preparation thereof) | | | |
| RN | 857890-33-6 CA | | | |
| CN | 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, hydrobromide (1:1) (CA INDEX NAME) | | | |



RN 857890-39-2 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]-
 methoxy-7-methoxy-, methanesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
 CMF C21 H19 Cl N4 O4



CM 2

CRN 75-75-2
 CMF C H4 O3 S



IT 857890-43-8P 857890-45-0P
 RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP (Physical process); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); RACT (Reactant or reagent); USES (Uses)
 (crystal of salt of 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino-phenoxy]-7-methoxy-6-quinolinicarboxamide or solvate thereof as neovascularization inhibitor, and preparation thereof)

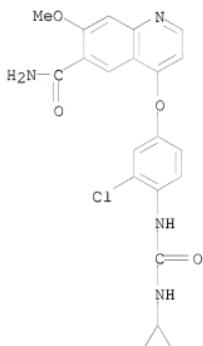
RN 857890-43-8 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, monomethanesulfonate, compd. with sulfonylbis[methane] (9CI) (CA INDEX NAME)

CM 1

CRN 417716-92-8

CMF C21 H19 Cl N4 O4



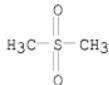
CM 2

CRN 75-75-2

CMF C H4 O3 S

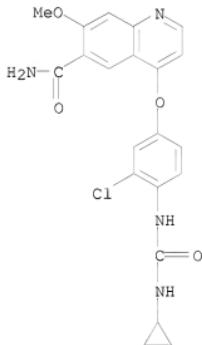


CM 3

CRN 67-71-0
CMF C2 H6 O2 S

RN 857890-45-0 CA
 CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
 henoxyl-7-methoxy-, acetate compd. with methanesulfonic acid (1:?:?) (CA
 INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4

CM 2

CRN 75-75-2
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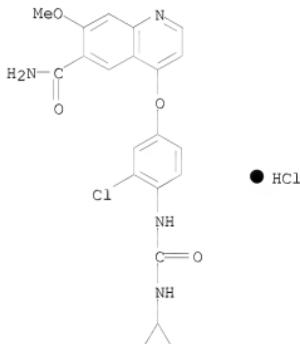
CM 3

CRN 64-19-7
CMF C2 H4 O2IT 857890-31-4P 857890-35-8P 857890-37-0P
857890-41-6P 857890-47-2P 857890-49-4P

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP (Physical process); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (crystal of salt of 4-[3-chloro-4-(cyclopropylamino)carbonyl]amino-phenoxy]-7-methoxy-6-quinolinelinecarboxamide or solvate thereof as neovascularization inhibitor, and preparation thereof)

RN 857890-31-4 CA

CN 6-Quinolinelinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, hydrochloride (1:1) (CA INDEX NAME)



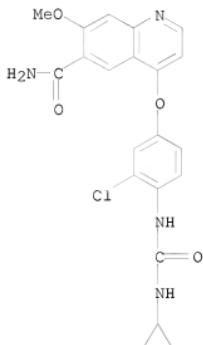
RN 857890-35-8 CA

CN 6-Quinolinelinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

CM 1

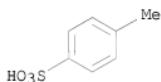
10/577531

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

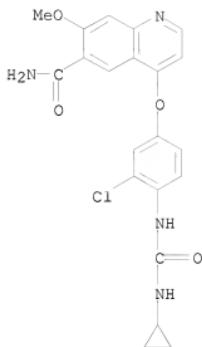
CRN 104-15-4
CMF C7 H8 O3 S



RN 857890-37-0 CA
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]oxy-, sulfate (1:1) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

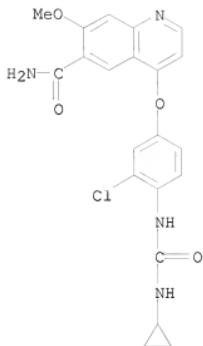
CRN 7664-93-9
CMF H₂ O₄ S



RN 857890-41-6 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p-henoxy]-7-methoxy-, methanesulfonate, hydrate (1:1:?) (CA INDEX NAME)

CM 1

CRN 417716-92-8
CMF C₂₁ H₁₉ Cl N₄ O₄



CM 2

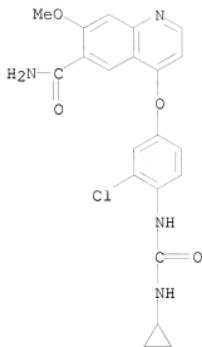
CRN 75-75-2
CMF C H4 O3 S



RN 857890-47-2 CA
CN Ethanesulfonic acid, compd. with 4-[3-chloro-4-
[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-6-
quinolinecarboxamide (1:1) (CA INDEX NAME)

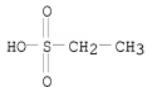
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CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

CRN 594-45-6
CMF C2 H6 O3 S

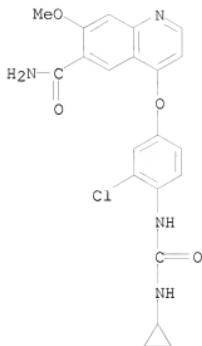


RN 857890-49-4 CA

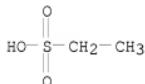
CN Ethanesulfonic acid, compd. with 4-[3-chloro-4-
[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy-6-
quinolinicarboxamide and sulfonylbis[methane] (1:1:?) (9CI) (CA INDEX
NAME)

CM 1

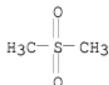
CRN 417716-92-8
CMF C21 H19 Cl N4 O4



CM 2

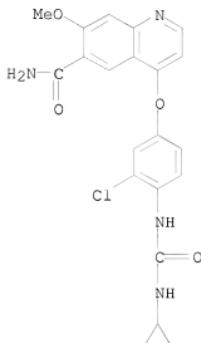
CRN 594-45-6
CMF C2 H6 O3 S

CM 3

CRN 67-71-0
CMF C2 H6 O2 S

IT 417716-92-8P, 4-[3-Chloro-4-(cyclopropylaminocarbonyl)amino-phenoxyl]-7-methoxy-6-quinolinicarboxamide
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (crystal of salt of 4-[3-chloro-4-(cyclopropylaminocarbonyl)amino-phenoxyl]-7-methoxy-6-quinolinicarboxamide or solvate thereof as neovascularization inhibitor, and preparation thereof)
 RN 417716-92-8 CA
 CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p

heenoxy]-7-methoxy- (CA INDEX NAME)

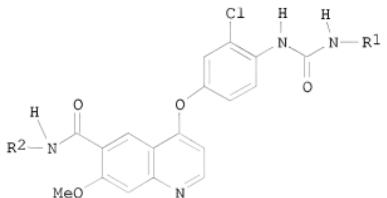


REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 19 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 142:481959 CA
 TITLE: Preparation of urea moiety-containing quinoliniccarboxamide derivatives
 INVENTOR(S): Naito, Toshihiko; Yoshizawa, Kazuhiro
 PATENT ASSIGNEE(S): Eisai Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 21 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2005044788 | A1 | 20050519 | WO 2004-JP16526 | 20041108 |
| W: AB, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1683785 | A1 | 20060726 | EP 2004-818213 | 20041108 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR, IS, YU | | | | |

| | | | |
|------------------------|--|------------------|------------|
| CN 1878751 | A 20061213 | CN 2004-80033071 | 20041108 |
| US 20070037849 | A1 20070215 | US 2006-577308 | 20060428 |
| IN 2006CN02045 | A 20070601 | IN 2006-CN2045 | 20060609 |
| PRIORITY APPLN. INFO.: | | JP 2003-381249 | A 20031111 |
| | | WO 2004-JP16526 | W 20041108 |
| OTHER SOURCE(S): | CASREACT 142:481959; MARPAT 142:481959 | | |
| GI | | | |

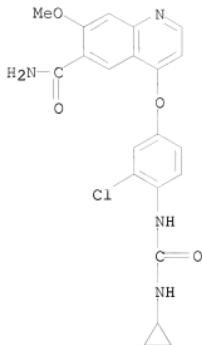


AB The title compds. I [wherein R1 is hydrogen, C1-6 alkyl, or C3-8 cycloalkyl; and R2 is hydrogen or methoxyl] are prepared by reaction of 4-amino-3-chlorophenol with aryl chloroformate, followed by reaction with an amine and reaction of the resulting urea derivative with a chloroquinoline derivative. I are useful in the treatment of diseases accompanied by abnormal proliferation of angiogenesis (no data). Thus, reaction of 4-amino-3-chlorophenol with Ph chloroformate, followed by reaction with cyclopropylamine and reaction of the resulting urea derivative with 7-methoxy-4-chloroquinoline-6-carboxamide, gave 4-(3-chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide.

IT 417716-92-8P
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (amination of aryl chloroformate or amination of aryl N-hydroxyphenylcarbamate)

RN 417716-92-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy-7-methoxy- (CA INDEX NAME)



REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 20 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 141:427993 CA
 TITLE: Polymorphous crystal of 4-(3-chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinicarboxamide and method for preparation thereof
 INVENTOR(S): Arimoto, Itaru; Yoshizawa, Kazuhiro; Kamada, Atsushi
 PATENT ASSIGNEE(S): Eisai Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 79 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|---|--------------------------------------|
| WO 2004101526 | A1 | 20041125 | WO 2004-JP5788 | 20040422 |
| W: AB, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| US 20070117842 | A1 | 20070524 | US 2006-553927 US 2003-464674P WO 2004-JP5788 | 20060630 P 20030422 W 20040422 |
| PRIORITY APPLN. INFO.: | | | | |

AB Disclosed are a polymorphous crystal (A) of 4-(3-chloro-4-

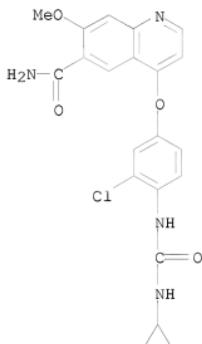
(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide (I) having a diffraction peak at a diffraction angle ($2\theta \pm 0.2^\circ$) of 15.75° in the powder X-ray diffractometry; and a polymorphous crystal (B) of I having a diffraction peak at a diffraction angle ($2\theta \pm 0.2^\circ$) of 21.75° in the powder X-ray diffractometry.

IT 417716-92-8P

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); PRP (Properties); PUR (Purification or recovery); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
 (preparation of 4-(3-chloro-4-(cyclopropylaminocarbonyl)aminophenoxy)-7-methoxy-6-quinolinecarboxamide polymorphous crystals)

RN 417716-92-8 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 21 OF 22 CA COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 141:289013 CA

TITLE: c-Kit kinase inhibitor

INVENTOR(S): Yamamoto, Yuji; Watanabe, Tatsuo; Okada, Masayuki; Tsuruoka, Akihiko

PATENT ASSIGNEE(S): Eisai Co., Ltd., Japan

SOURCE: PCT Int. Appl., 28 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|------|----------|-----------------|----------|
| WO 2004080462 | A1 | 20040923 | WO 2004-JP3087 | 20040310 |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,

CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
 GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
 LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
 NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
 TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
 BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
 ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
 SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
 TD, TG

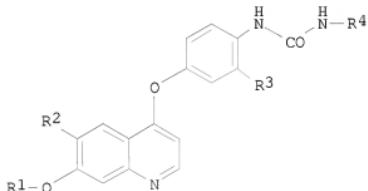
US 20040253205 A1 20041216 US 2004-797903 20040310

EP 1604665 A1 20051214 EP 2004-719054 20040310

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK

PRIORITY APPLN. INFO.: JP 2003-62823 A 20030310
 JP 2003-302803 A 20030827
 WO 2004-JP3087 W 20040310

OTHER SOURCE(S): MARPAT 141:289013
 GI



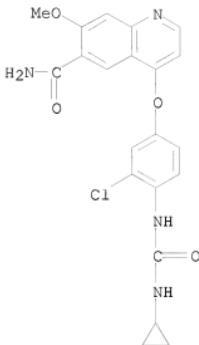
AB It is found out that a compound represented by the following general formula I (R1 = Me, etc.; R2 = cyano, etc.; R3 = H, etc.; and R4 = H, etc.) shows a potent c-Kit kinase inhibitory activity and suppresses the proliferation of cancer cells activated by c-Kit kinase both in vitro and in vivo. Thus, a novel anticancer agent showing a c-Kit kinase inhibitory activity is found out.

IT 417716-92-8, 4-(3-Chloro-4-((cyclopropylaminocarbonyl)amino)phenoxyl)-7-methoxy-6-quinolinecarboxamide

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (c-Kit kinase inhibitor)

RN 417716-92-8 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylaminocarbonyl)amino]phenoxy]-7-methoxy- (CA INDEX NAME)



REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

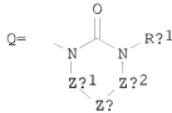
L4 ANSWER 22 OF 22 CA COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 136:340689 CA
 TITLE: Preparation of urea derivatives containing nitrogenous aromatic ring compounds as inhibitors of angiogenesis
 INVENTOR(S): Funahashi, Yasuhiro; Tsuruoka, Akihiko; Matsukura, Masayuki; Haneda, Toru; Fukuda, Yoshio; Kamata, Junichi; Takahashi, Keiko; Matsushima, Tomohiro; Miyazaki, Kazuki; Nomoto, Kenichi; Watanabe, Tatsuo; Obaishi, Hiroshi; Yamaguchi, Atsumi; Suzuki, Sachi; Nakamura, Katsuji; Mimura, Fusayo; Yamamoto, Yuji; Matsui, Junji; Matsui, Kenji; Yoshioka, Takako; Suzuki, Yasuyuki; Arimoto, Itaru
 PATENT ASSIGNEE(S): Eisai Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 699 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2002032872 | A1 | 20020425 | WO 2001-JP9221 | 20011019 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NC, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |

| | | | | |
|--|----|----------|------------------|-------------|
| CA 2426461 | A1 | 20020425 | CA 2001-2426461 | 20011019 |
| AU 2001095986 | A | 20020429 | AU 2001-95986 | 20011019 |
| HU 2003002603 | A2 | 20031128 | HU 2003-2603 | 20011019 |
| CN 1478078 | A | 20040225 | CN 2001-819710 | 20011019 |
| EP 1415987 | A1 | 20040506 | EP 2001-976786 | 20011019 |
| EP 1415987 | B1 | 20070228 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY, TR | | | | |
| EP 1506962 | A2 | 20050216 | EP 2004-25700 | 20011019 |
| EP 1506962 | A3 | 20050302 | | |
| EP 1506962 | B1 | 20080702 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY, TR | | | | |
| NZ 525324 | A | 20050324 | NZ 2001-525324 | 20011019 |
| JP 3712393 | B2 | 20051102 | JP 2002-536056 | 20011019 |
| RU 2264389 | C2 | 20051120 | RU 2003-114740 | 20011019 |
| AT 355275 | T | 20060315 | AT 2001-976786 | 20011019 |
| AU 2001295986 | B2 | 20060817 | AU 2001-295986 | 20011019 |
| EP 1777218 | A1 | 20070425 | EP 2006-23078 | 20011019 |
| R: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE, TR | | | | |
| CN 101024627 | A | 20070829 | CN 2007-10007096 | 20011019 |
| CN 101029022 | A | 20070905 | CN 2007-10007097 | 20011019 |
| ES 2282299 | T3 | 20071016 | ES 2001-976786 | 20011019 |
| AT 399766 | T | 20080715 | AT 2004-25700 | 20011019 |
| NO 2003001731 | A | 20030619 | NO 2003-1731 | 20030414 |
| MX 2003PA03362 | A | 20030801 | MX 2003-PA3362 | 20030415 |
| US 7253286 | B2 | 20070807 | US 2003-420466 | 20030418 |
| US 20040053908 | A1 | 20040318 | | |
| ZA 2003003567 | A | 20040810 | ZA 2003-3567 | 20030508 |
| JP 2005272474 | A | 20051006 | JP 2005-124034 | 20050421 |
| US 20060247259 | A1 | 20061102 | US 2005-293785 | 20051202 |
| US 20060160832 | A1 | 20060720 | US 2006-347749 | 20060203 |
| AU 2006203099 | A1 | 20060810 | AU 2006-203099 | 20060719 |
| AU 2006236039 | A1 | 20061207 | AU 2006-236039 | 20061116 |
| AU 2006236039 | B2 | 20080522 | | |
| NO 2007004657 | A | 20030619 | NO 2007-4657 | 20070912 |
| PRIORITY APPLN. INFO.: | | | JP 2000-320420 | A 20001020 |
| | | | JP 2000-386195 | A 20001220 |
| | | | JP 2001-46685 | A 20010222 |
| | | | AU 2001-295986 | A3 20011019 |
| | | | AU 2001-95986 | TO 20011019 |
| | | | CN 2001-819710 | A3 20011019 |
| | | | EP 2001-976786 | A3 20011019 |
| | | | JP 2002-536056 | A3 20011019 |
| | | | WO 2001-JP9221 | W 20011019 |
| | | | US 2003-420466 | A3 20030418 |
| | | | US 2005-293785 | A1 20051202 |

OTHER SOURCE(S):
GI

MARPAT 136:340689



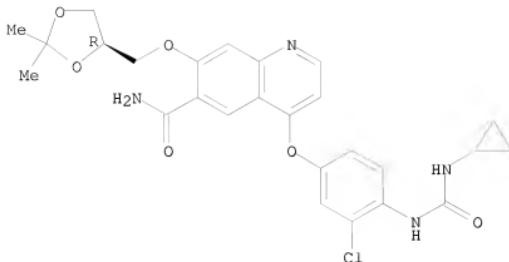
AB N-aryl or N-heteroarylurea derivs. represented by the general formula Ag-Xg-Yg-Tg1 or salts thereof, or hydrates of both [wherein Ag = (un)substituted C6-14 aryl or 5- to 14-membered heterocyclic group; Xg = single bond, O, S, Cl-6 alkylene, SO, SO₂, (un)substituted NH; Yg = (un)substituted C6-14 aryl, 5- to 14-membered heterocyclic group, Cl-8 alkyl, C3-8 alicyclic hydrocarbyl, C6-14 aryl-Cl-6 alkyl, 5- to 14-membered heteroaryl-Cl-6 alkyl, $(\text{CH}_2)_g\text{SO}_2$ ($g = 1-8$), $(\text{CH}_2)_f\text{A}-\text{CH}(\text{CH}_2)\text{fb}$ ($f, b = 0, 1, 2, 3$), etc.; and Tg1 = a group of the general formula -Eg-CO-NRg1(Zg) or Q; wherein Eg = a single bond, (un)substituted NH; Rg1 = H, (un)substituted Cl-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, C3-8 aliphatic hydrocarbyl, etc.; Zg = Cl-8 alkyl, C3-8 alicyclic hydrocarbyl, C6-14 aryl, etc.; Zg1, Zg2 = (a) a single bond, (b) Cl-6 alkylene optionally having ≥ 1 atoms selected from O, S, and N in the middle or the terminus of the chain and optionally substituted with o xo, (c) (un)substituted C2-6 alkenyl] are prepared. These compds. are also inhibitors of vascular endothelial growth factor receptor kinase (VEGFR2 kinase) and are useful as antitumor agents against hemangioma, pancreatic cancer, stomach cancer, colon cancer, breast cancer, prostate cancer, lung cancer, brain tumor, leukemia, or ovarian cancer, as cancer metastasis inhibitors, and for the treatment of retina neovascularization, diabetic retinopathy, atherosclerosis, or inflammatory diseases such as osteoarthritis, rheumatoid arthritis, psoriasis, or delayed hypersensitivity. Thus, to solution of 334 mg 4-[6-(4-benzyloxyphenyl)-7-(2-trimethylsilylethoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy]-2-chlorophenylamine in 4 mL DMF were added 0.066 mL pyridine and 0.102 mL Ph chlorocarbonate and stirred at room temperature for 2.5 h to give 330 mg N-[4-(4-benzyloxyphenyl)-7-(2-trimethylsilylethoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy]-2-chlorophenyl-N'-cyclopropylurea which (260 mg) was hydrogenolyzed over platinum oxide in ethanol overnight to give 160 mg N-[4-(6-(4-hydroxyphenyl)-7-(2-trimethylsilylethoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy)-2-chlorophenyl]-N'-cyclopropylurea (I). I showed IC₅₀ of 0.02 nM for inhibiting the vascular endothelial growth factor (VEGF)-stimulated sandwich tube formation in vascular endothelial cell.

IT 417717-12-5P 417717-13-6P
RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (preparation of urea derivs. containing nitrogenous aromatic ring compds. as angiogenesis inhibitors for prevention or treatment of diseases)

RN 417717-12-5 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(4R)-2,2-dimethyl-1,3-dioxolan-4-yl]methoxy- (CA INDEX NAME)

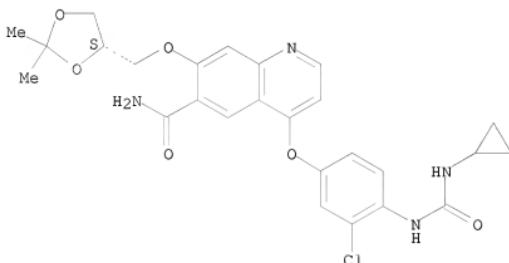
Absolute stereochemistry.



RN 417717-13-6 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(4S)-2-dimethyl-1,3-dioxolan-4-yl]methoxy- (CA INDEX NAME)

Absolute stereochemistry.



IT 417716-92-8P 417717-03-4P 417717-05-6P

417717-06-7P 417717-07-8P 417717-08-9P

417717-09-0P 417717-10-3P 417717-11-4P

417717-14-7P 417717-15-8P 417717-16-9P

417717-17-0P 417717-18-1P 417717-19-2P

417719-50-7P 417719-56-3P 417719-57-4P

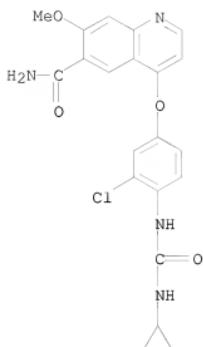
417719-77-8P 417720-06-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

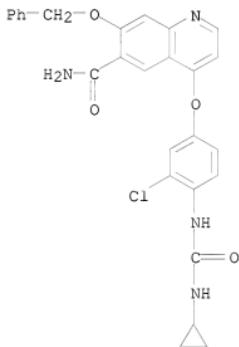
(preparation of urea derivs. containing nitrogenous aromatic ring compds. as angiogenesis inhibitors for prevention or treatment of diseases)

RN 417716-92-8 CA

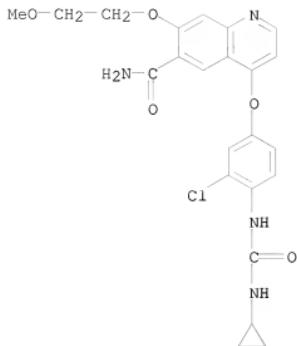
CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-methoxy- (CA INDEX NAME)



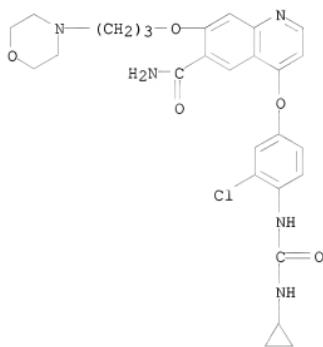
RN 417717-03-4 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-(phenylmethoxy)- (CA INDEX NAME)



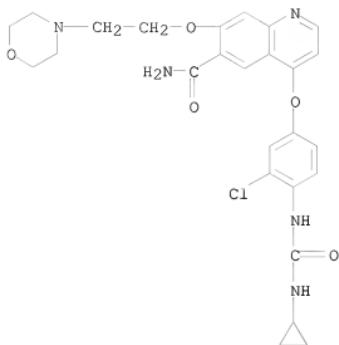
RN 417717-05-6 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]p
henoxy]-7-(2-methoxyethoxy)- (CA INDEX NAME)



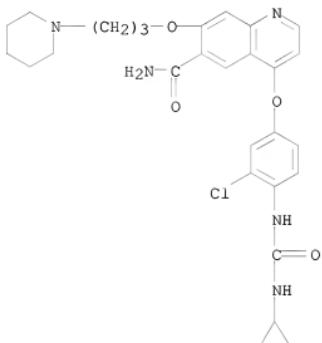
RN 417717-06-7 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[3-(4-morpholinyl)propoxy]- (CA INDEX NAME)



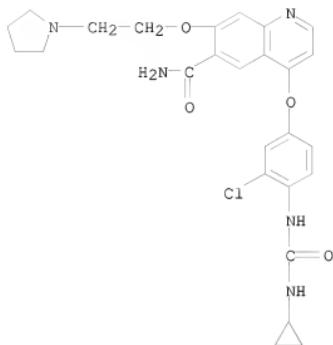
RN 417717-07-8 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(4-morpholinyl)ethoxy]- (CA INDEX NAME)



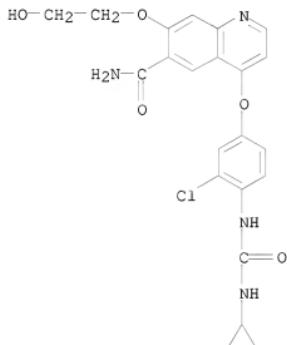
RN 417717-08-9 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[3-(1-piperidinyl)propoxy]- (CA INDEX NAME)



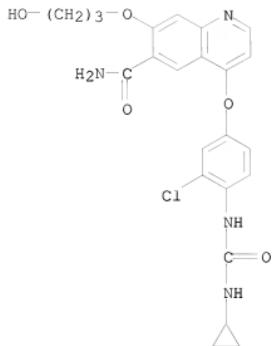
RN 417717-09-0 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[2-(1-pyrrolidinyl)ethoxy]- (CA INDEX NAME)



RN 417717-10-3 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-hydroxyethoxy)- (CA INDEX NAME)

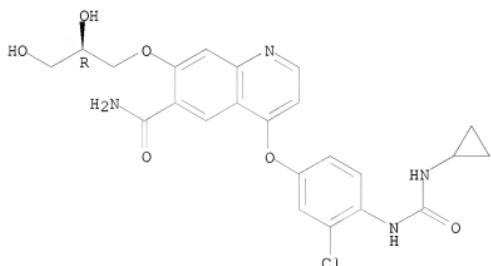


RN 417717-11-4 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(3-hydroxypropoxy)- (CA INDEX NAME)



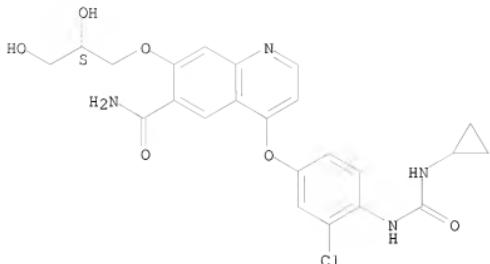
RN 417717-14-7 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



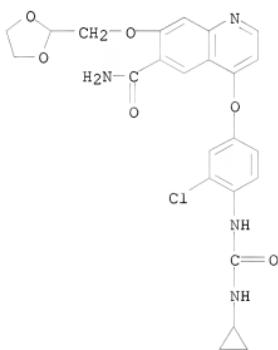
RN 417717-15-8 CA
CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2S)-2,3-dihydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



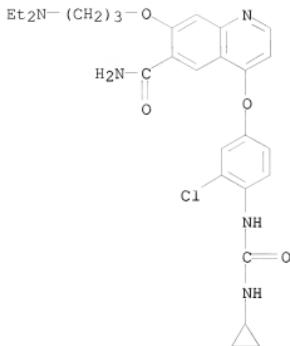
RN 417717-16-9 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(1,3-dioxolan-2-ylmethoxy)- (CA INDEX NAME)



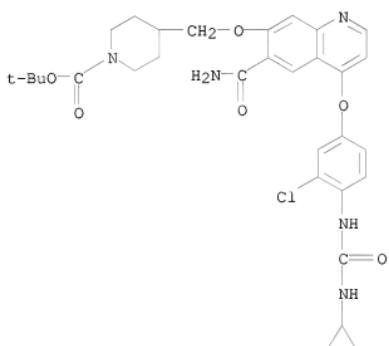
RN 417717-17-0 CA

CN 6-Quinolinecarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[3-(diethylamino)propoxy]- (CA INDEX NAME)



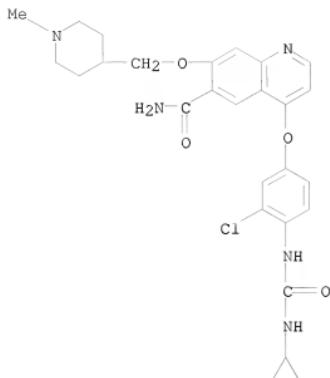
RN 417717-18-1 CA

CN 1-Piperidinecarboxylic acid, 4-[(6-(aminocarbonyl)-4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy)-7-quinolinyl]oxy)methyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)



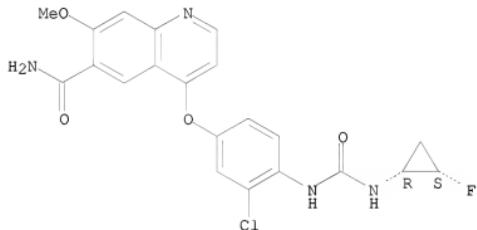
RN 417717-19-2 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(1-methyl-4-piperidinyl)methoxy]- (CA INDEX NAME)

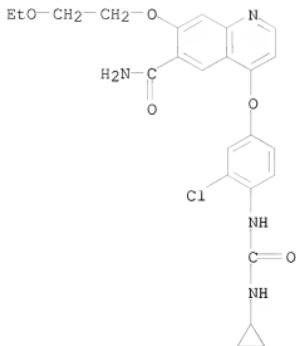


RN 417719-50-7 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[[[(1R,2S)-2-fluorocyclopropyl]amino]carbonyl]amino]phenoxy-, rel- (CA INDEX NAME)

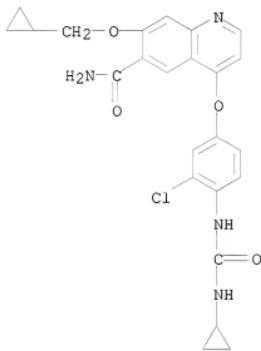
Relative stereochemistry.



RN 417719-56-3 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(2-ethoxyethoxy)- (CA INDEX NAME)

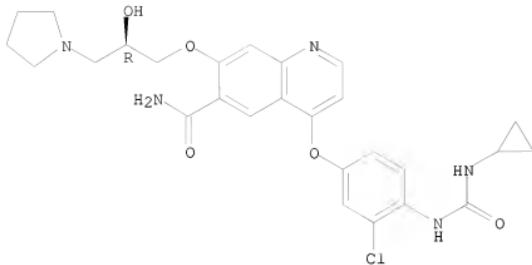


RN 417719-57-4 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(cyclopropylmethoxy)- (CA INDEX NAME)



RN 417719-77-8 CA
CN 6-Quinolinicarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-{(2R)-2-hydroxy-3-(1-pyrrolidinyl)propoxy}- (CA INDEX NAME)

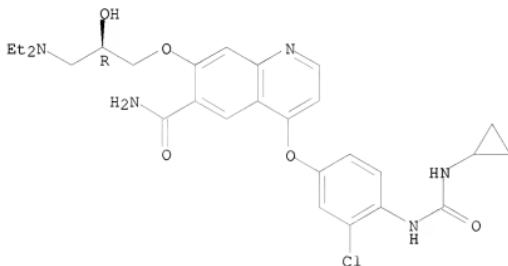
Absolute stereochemistry.



RN 417720-06-0 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-[(2R)-3-(diethylamino)-2-hydroxypropoxy]- (CA INDEX NAME)

Absolute stereochemistry.



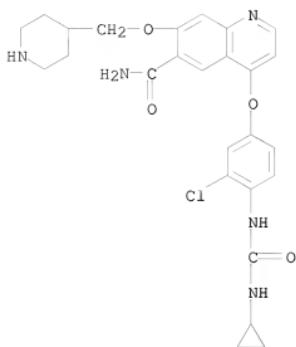
IT 417724-98-2P, 4-(3-Chloro-4-((cyclopropylamino)carbonyl)amino)phenoxy)-7-((4-piperidyl)methoxy)-6-quinoliniccarboxamide

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of urea derivs. containing nitrogenous aromatic ring compds. as angiogenesis inhibitors for prevention or treatment of diseases)

RN 417724-98-2 CA

CN 6-Quinoliniccarboxamide, 4-[3-chloro-4-[(cyclopropylamino)carbonyl]amino]phenoxy]-7-(4-piperidinylmethoxy)- (CA INDEX NAME)



REFERENCE COUNT:

17

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